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(57) Abstract: Nucleotide sequences are isolated from *Arabidopsis thaliana* that code for proteins essential for plant growth and developement. The essentially of the proteins may be exploited by recombinantly expressing the proteins and using them in screening assays to identify compounds that interact with or inhibit the proteins and are therefore potential herbicides.



NUCLEIC ACID MOLECULES ENCODING PROTEINS ESSENTIAL FOR PLANT GROWTH AND DEVELOPMENT AND USES THEREOF

The present invention pertains to nucleic acid molecules isolated from Arabidopsis thaliana comprising nucleotide sequences that encode proteins essential for plant growth and development. The invention particularly relates to methods of using these proteins as herbicide targets, based on this essentiality.

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The use of herbicides to control undesirable vegetation such as weeds in crop fields has become almost a universal practice. The herbicide market exceeds 15 billion dollars annually. Despite this extensive use, weed control remains a significant and costly problem for farmers.

Effective use of herbicides requires sound management. For instance, the time and method of application and stage of weed plant development are critical to achieving good weed control with herbicides. Because various weed species are resistant to herbicides, the production of effective new herbicides becomes increasingly important. New herbicides can now be discovered using high-throughput screens that implement recombinant DNA technology. Metabolic enzymes found to be essential to plant growth and development can be recombinantly produced through standard molecular biological techniques and utilized as herbicide targets in screens for novel inhibitors of the enzyme activity. More generally, any essential plant protein can be used to screen for inhibitors of its activity. The novel inhibitors discovered through such screens may then be used as herbicides to control undesirable vegetation.

In view of the above, there remain persistent and ongoing problems with unwanted or detrimental vegetation growth (e.g. weeds). Furthermore, as the population continues to grow, there will be increasing food shortages. Therefore, there exists a long felt, yet unfulfilled need, to find new, effective, and economic herbicides.

In view of these needs, it is an object of the invention to provide nucleic acid molecules from Arabidopsis thaliana comprising nucleotide sequences that encode proteins essential for plant growth and development. It is another object to provide the essential proteins encoded by these essential nucleotide sequences for assay development to identify

inhibitory compounds with herbicidal activity. It is still another object of the present invention to provide an effective and beneficial method for identifying new or improved herbicides using the essential proteins of the invention.

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In furtherance of these and other objects, the present invention provides nucleic acid molecules isolated from Arabidopsis thaliana comprising nucleotide sequences that encode proteins essential for plant viability. Genetic results show that when any of the nucleotide sequences of the invention are mutated in Arabidopsis thaliana, the resulting phenotype is embryo or seedling lethal in the homozygous state. In particular, by using Ac/Ds transposon or T-DNA-mediated mutagenesis, the inventors of the present invention are the first to demonstrate that the activity of each protein of the present invention is essential for plant growth in Arabidopsis thaliana.

This knowledge is exploited to provide novel herbicide modes of action. The critical role in plant growth of the proteins encoded by each of the nucleotide sequences of the invention implies that chemicals that inhibit the function of any one of these proteins in plants are likely to have detrimental effects on plants and are potentially good herbicide candidates. Thus, the proteins encoded by the essential nucleotide sequences provide the bases for assays designed to easily and rapidly identify novel herbicides.

The present invention therefore provides methods of using a purified protein encoded by any one of the nucleotide sequences described below to identify inhibitors thereof, which can then be used as herbicides to suppress the growth of undesirable vegetation, e.g. in fields where crops are grown, particularly agronomically important crops such as maize and other cereal crops such as wheat, oats, rye, sorghum, rice, barley, millet, turf and forage grasses, and the like, as well as cotton, sugar cane, sugar beet, oilseed rape, and soybeans.

Disclosed herein are nucleic acid molecules isolated from *Arabidopsis thaliana*. In one embodiment, the present invention provides an isolated nucleic acid molecule comprising a nucleotide sequence, the complement of which hybridizes under stringent conditions to a sequence selected from the group consisting of the odd numbered SEQ ID NOs:1-95. In another embodiment, the present invention provides an isolated nucleic acid molecule comprising a nucleotide sequence that encodes a protein comprising an amino acid sequence having at least 60%, preferably 70%, more preferably 80%, still more preferably 90%, even more preferably 95%, and most preferably 99-100% sequence identity to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96.

The present invention also provides a chimeric construct comprising a promoter operatively linked to a nucleic acid molecule according to the present invention, wherein the promoter is preferably functional in a eukaryote, wherein the promoter is preferably heterologous to the nucleic acid molecule. The present invention further provides a recombinant vector comprising a chimeric construct according to the present invention, wherein said vector is capable of being stably transformed into a host cell. The present invention still further provides a host cell comprising a nucleic acid molecule according to the present invention, wherein said nucleic acid molecule is preferably expressible in the cell. The host cell is preferably selected from the group consisting of a plant cell, a yeast cell, an insect cell, and a prokaryotic cell. The present invention additionally provides a plant or seed comprising a plant cell according to the present invention.

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The present invention also provides proteins essential for plant growth in *Arabidopsis thaliana*. In one embodiment, the present invention provides an isolated protein comprising an amino acid sequence having at least 60%, preferably 70%, more preferably 80%, still more preferably 90%, even more preferably 95%, and most preferably 99-100% sequence identity to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96. In accordance with another embodiment, the present invention also relates to the recombinant production of proteins of the invention and methods of using the proteins of the invention in assays for identifying compounds that interact with the protein.

According to another aspect, the present invention provides a method of identifying a herbicidal compound, comprising: (a) combining a polypeptide comprising an amino acid sequence at least 90% identical to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96 with a compound to be tested for the ability to bind to said polypeptide, under conditions conducive to binding; (b) selecting a compound identified in (a) that binds to said polypeptide; (c) applying a compound selected in (b) to a plant to test for herbicidal activity; and (d) selecting a compound identified in (c) that has herbicidal activity. Preferably, the polypeptide comprises an amino acid sequence at least 95% identical to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96. More preferably, the polypeptide comprises an amino acid sequence at least 99% identical to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96. Most preferably, the polypeptide comprises an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96. Most preferably, the polypeptide comprises an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96.

NOs:2-96. The present invention also provides a method for killing or inhibiting the growth or viability of a plant, comprising applying to the plant a herbicidal compound identified according to this method.

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According to yet another aspect, the present invention provides a method of identifying a herbicidal compound, comprising: (a) combining a polypeptide comprising an amino acid sequence at least 90% identical to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96 with a compound to be tested for the ability to inhibit the activity of said polypeptide, under conditions conducive to inhibition; (b) selecting a compound identified in (a) that inhibits the activity of said polypeptide; (c) applying a compound selected in (b) to a plant to test for herbicidal activity; and (d) selecting a compound identified in (c) that has herbicidal activity. Preferably, the polypeptide comprises an amino acid sequence at least 95% identical to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96. More preferably, the polypeptide comprises an amino acid sequence at least 99% identical to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96. Most preferably, the polypeptide comprises an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96. The present invention also provides a method for killing or inhibiting the growth or viability of a plant, comprising applying to the plant a herbicidal compound identified according to this method.

The present invention still further provides a method for killing or inhibiting the growth or viability of a plant, comprising inhibiting expression in said plant of a protein having at least 60%, preferably 70%, more preferably 80%, still more preferably 90%, even more preferably 95%, and most preferably 99-100% sequence identity to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96.

Other objects and advantages of the present invention will become apparent to those skilled in the art and from a study of the following description of the invention and non-limiting examples. The entire contents of all publications mentioned herein are hereby incorporated by reference.

BRIEF DESCRIPTION OF THE SEQUENCES IN THE SEQUENCE LISTING
Odd numbered SEQ ID NOs:1-95 are nucleotide sequences isolated from
Arabidopsis thaliana that are more fully described in Table 5 below.

Even numbered SEQ ID NOs:2-96 are protein sequences encoded by the immediately preceding nucleotide sequence, e.g., SEQ ID NO:2 is the protein encoded by the nucleotide sequence of SEQ ID NO:1, SEQ ID NO:4 is the protein encoded by the nucleotide sequence of SEQ ID NO:3, etc.

SEQ ID NOs:101-125 are PCR primers.

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DEFINITIONS

For clarity, certain terms used in the specification are defined and presented as follows:

"Associated with / operatively linked" refer to two nucleic acid sequences that are related physically or functionally. For example, a promoter or regulatory DNA sequence is said to be "associated with" a DNA sequence that codes for an RNA or a protein if the two sequences are operatively linked, or situated such that the regulator DNA sequence will affect the expression level of the coding or structural DNA sequence.

A "chimeric construct" is a recombinant nucleic acid sequence in which a promoter or regulatory nucleic acid sequence is operatively linked to, or associated with, a nucleic acid sequence that codes for an mRNA or which is expressed as a protein, such that the regulatory nucleic acid sequence is able to regulate transcription or expression of the associated nucleic acid sequence. The regulatory nucleic acid sequence of the chimeric construct is not normally operatively linked to the associated nucleic acid sequence as found in nature.

Co-factor: natural reactant, such as an organic molecule or a metal ion, required in an enzyme-catalyzed reaction. A co-factor is *e.g.* NAD(P), riboflavin (including FAD and FMN), folate, molybdopterin, thiamin, biotin, lipoic acid, pantothenic acid and coenzyme A, S-adenosylmethionine, pyridoxal phosphate, ubiquinone, menaquinone. Optionally, a co-factor can be regenerated and reused.

A "coding sequence" is a nucleic acid sequence that is transcribed into RNA such as mRNA, rRNA, tRNA, snRNA, sense RNA or antisense RNA. Preferably the RNA is then translated in an organism to produce a protein.

Complementary: "complementary" refers to two nucleotide sequences that comprise antiparallel nucleotide sequences capable of pairing with one another upon formation of hydrogen bonds between the complementary base residues in the antiparallel nucleotide sequences.

Enzyme activity: means herein the ability of an enzyme to catalyze the conversion of a substrate into a product. A substrate for the enzyme comprises the natural substrate of the enzyme but also comprises analogues of the natural substrate, which can also be converted, by the enzyme into a product or into an analogue of a product. The activity of the enzyme is measured for example by determining the amount of product in the reaction after a certain period of time, or by determining the amount of substrate remaining in the reaction mixture after a certain period of time. The activity of the enzyme is also measured by determining the amount of an unused co-factor of the reaction remaining in the reaction mixture after a certain period of time or by determining the amount of used co-factor in the reaction mixture after a certain period of time. The activity of the enzyme is also measured by determining the amount of a donor of free energy or energy-rich molecule (e.g. ATP, phosphoenolpyruvate, acetyl phosphate or phosphocreatine) remaining in the reaction mixture after a certain period of time or by determining the amount of a used donor of free energy or energy-rich molecule (e.g. ADP, pyruvate, acetate or creatine) in the reaction mixture after a certain period of time.

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Essential: an "essential" Arabidopsis thaliana nucleotide sequence is a nucleotide sequence encoding a protein such as e.g. a biosynthetic enzyme, receptor, signal transduction protein, structural gene product, or transport protein that is essential to the growth or survival of the plant.

Expression Cassette: "Expression cassette" as used herein means a nucleic acid molecule capable of directing expression of a particular nucleotide sequence in an appropriate host cell, comprising a promoter operatively linked to the nucleotide sequence of interest which is operatively linked to termination signals. It also typically comprises sequences required for proper translation of the nucleotide sequence. The coding region usually codes for a protein of interest but may also code for a functional RNA of interest, for example antisense RNA or a nontranslated RNA, in the sense or antisense direction. The expression cassette comprising the nucleotide sequence of interest may be chimeric, meaning that at least one of its components is heterologous with respect to at least one of its other components. The expression cassette may also be one that is naturally occurring but has been obtained in a recombinant form useful for heterologous expression. Typically, however, the expression cassette is heterologous with respect to the host, i.e., the particular DNA sequence of the expression cassette does not occur naturally in the host cell and must have been introduced into the host cell or an ancestor of the host cell by a transformation event. The expression of

the nucleotide sequence in the expression cassette may be under the control of a constitutive promoter or of an inducible promoter that initiates transcription only when the host cell is exposed to some particular external stimulus. In the case of a multicellular organism, such as a plant, the promoter can also be specific to a particular tissue or organ or stage of development.

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Gene: the term "gene" is used broadly to refer to any segment of DNA associated with a biological function. Thus, genes include coding sequences and/or the regulatory sequences required for their expression. Genes also include nonexpressed DNA segments that, for example, form recognition sequences for other proteins. Genes can be obtained from a variety of sources, including cloning from a source of interest or synthesizing from known or predicted sequence information, and may include sequences designed to have desired parameters.

herein to refer to a nucleic acid sequence (e.g. a DNA sequence) or a gene, refer to a sequence that originates from a source foreign to the particular host cell or, if from the same source, is modified from its original form. Thus, a heterologous gene in a host cell includes a gene that is endogenous to the particular host cell but has been modified through, for example, the use of DNA shuffling. The terms also include non-naturally occurring multiple copies of a naturally occurring DNA sequence. Thus, the terms refer to a DNA segment that is foreign or heterologous to the cell, or homologous to the cell but in a position within the host cell nucleic acid in which the element is not ordinarily found. Exogenous DNA segments are expressed to yield exogenous polypeptides.

A "homologous" nucleic acid (e.g. DNA) sequence is a nucleic acid (e.g. DNA) sequence naturally associated with a host cell into which it is introduced.

Hybridization: The phrase "hybridizing specifically to" refers to the binding, duplexing, or hybridizing of a molecule only to a particular nucleotide sequence under stringent conditions when that sequence is present in a complex mixture (e.g., total cellular) DNA or RNA. "Bind(s) substantially" refers to complementary hybridization between a probe nucleic acid and a target nucleic acid and embraces minor mismatches that can be accommodated by reducing the stringency of the hybridization media to achieve the desired detection of the target nucleic acid sequence.

Inhibitor: a chemical substance that inactivates the enzymatic activity of a protein such as a biosynthetic enzyme, receptor, signal transduction protein, structural gene product, or transport protein. The term "herbicide" (or "herbicidal compound") is used herein to define an inhibitor applied to a plant at any stage of development, whereby the herbicide inhibits the growth of the plant or kills the plant.

Interaction: quality or state of mutual action such that the effectiveness or toxicity of one protein or compound on another protein is inhibitory (antagonists) or enhancing (agonists).

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A nucleic acid sequence is "isocoding with" a reference nucleic acid sequence when the nucleic acid sequence encodes a polypeptide having the same amino acid sequence as the polypeptide encoded by the reference nucleic acid sequence.

Isogenic: plants that are genetically identical, except that they may differ by the presence or absence of a heterologous DNA sequence.

Isolated: in the context of the present invention, an isolated DNA molecule or an isolated enzyme is a DNA molecule or enzyme that, by the hand of man, exists apart from its native environment and is therefore not a product of nature. An isolated DNA molecule or enzyme may exist in a purified form or may exist in a non-native environment such as, for example, in a transgenic host cell.

Mature protein: protein from which the transit peptide, signal peptide, and/or propeptide portions have been removed.

Minimal Promoter: the smallest piece of a promoter, such as a TATA element, that can support any transcription. A minimal promoter typically has greatly reduced promoter activity in the absence of upstream activation. In the presence of a suitable transcription factor, the minimal promoter functions to permit transcription.

Modified Enzyme Activity: enzyme activity different from that which naturally occurs in a plant (i.e. enzyme activity that occurs naturally in the absence of direct or indirect manipulation of such activity by man), which is tolerant to inhibitors that inhibit the naturally occurring enzyme activity.

Native: refers to a gene that is present in the genome of an untransformed plant cell.

Naturally occurring: the term "naturally occurring" is used to describe an object that can be found in nature as distinct from being artificially produced by man. For example, a protein or nucleotide sequence present in an organism (including a virus), which can be

isolated from a source in nature and which has not been intentionally modified by man in the laboratory, is naturally occurring.

Nucleic acid: the term "nucleic acid" refers to deoxyribonucleotides or ribonucleotides and polymers thereof in either single- or double-stranded form. Unless specifically limited, the term encompasses nucleic acids containing known analogues of natural nucleotides which have similar binding properties as the reference nucleic acid and are metabolized in a manner similar to naturally occurring nucleotides. Unless otherwise indicated, a particular nucleic acid sequence also implicitly encompasses conservatively modified variants thereof (e.g. degenerate codon substitutions) and complementary sequences and as well as the sequence explicitly indicated. Specifically, degenerate codon substitutions may be achieved by generating sequences in which the third position of one or more selected (or all) codons is substituted with mixed-base and/or deoxyinosine residues (Batzer et al., Nucleic Acid Res. 19: 5081 (1991); Ohtsuka et al., J. Biol. Chem. 260: 2605-2608 (1985); Rossolini et al., Mol. Cell. Probes 8: 91-98 (1994)). The terms "nucleic acid" or "nucleic acid sequence" may also be used interchangeably with gene, cDNA, and mRNA encoded by a gene.

"ORF" means open reading frame.

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Percent identity: the phrases "percent identical" or "percent identical," in the context of two nucleic acid or protein sequences, refers to two or more sequences or subsequences that have for example 60%, preferably 70%, more preferably 80%, still more preferably 90%, even more preferably 95%, and most preferably at least 99% nucleotide or amino acid residue identity, when compared and aligned for maximum correspondence, as measured using one of the following sequence comparison algorithms or by visual inspection. Preferably, the percent identity exists over a region of the sequences that is at least about 50 residues in length, more preferably over a region of at least about 100 residues, and most preferably the percent identity exists over at least about 150 residues. In an especially preferred embodiment, the percent identity exists over the entire length of the coding regions.

For sequence comparison, typically one sequence acts as a reference sequence to which test sequences are compared. When using a sequence comparison algorithm, test and reference sequences are input into a computer, subsequence coordinates are designated if necessary, and sequence algorithm program parameters are designated. The sequence comparison algorithm then calculates the percent sequence identity for the test sequence(s) relative to the reference sequence, based on the designated program parameters.

Optimal alignment of sequences for comparison can be conducted, e.g., by the local homology algorithm of Smith & Waterman, Adv. Appl. Math. 2: 482 (1981), by the homology alignment algorithm of Needleman & Wunsch, J. Mol. Biol. 48: 443 (1970), by the search for similarity method of Pearson & Lipman, Proc. Nat'l. Acad. Sci. USA 85: 2444 (1988), by computerized implementations of these algorithms (GAP, BESTFIT, FASTA, and TFASTA in the Wisconsin Genetics Software Package, Genetics Computer Group, 575 Science Dr., Madison, WI), or by visual inspection (see generally, Ausubel et al., infra).

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One example of an algorithm that is suitable for determining percent sequence identity and sequence similarity is the BLAST algorithm, which is described in Altschul et al., J. Mol. Biol. 215: 403-410 (1990). Software for performing BLAST analyses is publicly available through the National Center for Biotechnology Information (http://www.ncbi.nlm.nih.gov/). This algorithm involves first identifying high scoring sequence pairs (HSPs) by identifying short words of length W in the query sequence, which either match or satisfy some positive-valued threshold score T when aligned with a word of the same length in a database sequence. T is referred to as the neighborhood word score threshold (Altschul et al., 1990). These initial neighborhood word hits act as seeds for initiating searches to find longer HSPs containing them. The word hits are then extended in both directions along each sequence for as far as the cumulative alignment score can be increased. Cumulative scores are calculated using, for nucleotide sequences, the parameters M (reward score for a pair of matching residues; always > 0) and N (penalty score for mismatching residues; always < 0). For amino acid sequences, a scoring matrix is used to calculate the cumulative score. Extension of the word hits in each direction are halted when the cumulative alignment score falls off by the quantity X from its maximum achieved value, the cumulative score goes to zero or below due to the accumulation of one or more negative-scoring residue alignments, or the end of either sequence is reached. The BLAST algorithm parameters W, T, and X determine the sensitivity and speed of the alignment. The BLASTN program (for nucleotide sequences) uses as defaults a wordlength (W) of 11, an expectation (E) of 10, a cutoff of 100, M=5, N=-4, and a comparison of both strands. For amino acid sequences, the BLASTP program uses as defaults a wordlength (W) of 3, an expectation (E) of 10, and the BLOSUM62 scoring matrix (see Henikoff & Henikoff, Proc. Natl. Acad. Sci. USA 89: 10915 (1989)).

In addition to calculating percent sequence identity, the BLAST algorithm also performs a statistical analysis of the similarity between two sequences (see, e.g., Karlin &

Altschul, *Proc. Nat'l. Acad. Sci. USA* 90: 5873-5787 (1993)). One measure of similarity provided by the BLAST algorithm is the smallest sum probability (P(N)), which provides an indication of the probability by which a match between two nucleotide or amino acid sequences would occur by chance. For example, a test nucleic acid sequence is considered similar to a reference sequence if the smallest sum probability in a comparison of the test nucleic acid sequence to the reference nucleic acid sequence is less than about 0.1, more preferably less than about 0.01, and most preferably less than about 0.001.

Pre-protein: protein that is normally targeted to a cellular organelle, such as a chloroplast, and still comprises its native transit peptide.

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Purified: the term "purified," when applied to a nucleic acid or protein, denotes that the nucleic acid or protein is essentially free of other cellular components with which it is associated in the natural state. It is preferably in a homogeneous state although it can be in either a dry or aqueous solution. Purity and homogeneity are typically determined using analytical chemistry techniques such as polyacrylamide gel electrophoresis or high performance liquid chromatography. A protein that is the predominant species present in a preparation is substantially purified. The term "purified" denotes that a nucleic acid or protein gives rise to essentially one band in an electrophoretic gel. Particularly, it means that the nucleic acid or protein is at least about 50% pure, more preferably at least about 85% pure, and most preferably at least about 99% pure.

Two nucleic acids are "recombined" when sequences from each of the two nucleic acids are combined in a progeny nucleic acid. Two sequences are "directly" recombined when both of the nucleic acids are substrates for recombination. Two sequences are "indirectly recombined" when the sequences are recombined using an intermediate such as a cross-over oligonucleotide. For indirect recombination, no more than one of the sequences is an actual substrate for recombination, and in some cases, neither sequence is a substrate for recombination.

"Regulatory elements" refer to sequences involved in controlling the expression of a nucleotide sequence. Regulatory elements comprise a promoter operatively linked to the nucleotide sequence of interest and termination signals. They also typically encompass sequences required for proper translation of the nucleotide sequence.

Significant Increase: an increase in enzymatic activity that is larger than the margin of error inherent in the measurement technique, preferably an increase by about 2-fold or greater

of the activity of the wild-type enzyme in the presence of the inhibitor, more preferably an increase by about 5-fold or greater, and most preferably an increase by about 10-fold or greater.

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Significantly less: means that the amount of a product of an enzymatic reaction is reduced by more than the margin of error inherent in the measurement technique, preferably a decrease by about 2-fold or greater of the activity of the wild-type enzyme in the absence of the inhibitor, more preferably an decrease by about 5-fold or greater, and most preferably an decrease by about 10-fold or greater.

Specific Binding/Immunological Cross-Reactivity: An indication that two nucleic acid sequences or proteins are substantially identical is that the protein encoded by the first nucleic acid is immunologically cross reactive with, or specifically binds to, the protein encoded by the second nucleic acid. Thus, a protein is typically substantially identical to a second protein, for example, where the two proteins differ only by conservative substitutions. The phrase "specifically (or selectively) binds to an antibody," or "specifically (or selectively) immunoreactive with," when referring to a protein or peptide, refers to a binding reaction which is determinative of the presence of the protein in the presence of a heterogeneous population of proteins and other biologics. Thus, under designated immunoassay conditions. the specified antibodies bind to a particular protein and do not bind in a significant amount to other proteins present in the sample. Specific binding to an antibody under such conditions may require an antibody that is selected for its specificity for a particular protein. For example, antibodies raised to the protein with the amino acid sequence encoded by any of the nucleic acid sequences of the invention can be selected to obtain antibodies specifically immunoreactive with that protein and not with other proteins except for polymorphic variants. A variety of immunoassay formats may be used to select antibodies specifically immunoreactive with a particular protein. For example, solid-phase ELISA immunoassays. Western blots, or immunohistochemistry are routinely used to select monoclonal antibodies specifically immunoreactive with a protein. See Harlow and Lane (1988) Antibodies, A Laboratory Manual, Cold Spring Harbor Publications, New York "Harlow and Lane"), for a description of immunoassay formats and conditions that can be used to determine specific immunoreactivity. Typically a specific or selective reaction will be at least twice background signal or noise and more typically more than 10 to 100 times background.

"Stringent hybridization conditions" and "stringent hybridization wash conditions" in the context of nucleic acid hybridization experiments such as Southern and Northern hybridizations are sequence dependent, and are different under different environmental parameters. Longer sequences hybridize specifically at higher temperatures. An extensive guide to the hybridization of nucleic acids is found in Tijssen (1993) *Laboratory Techniques in Biochemistry and Molecular Biology-Hybridization with Nucleic* Acid Probes part I chapter 2 "Overview of principles of hybridization and the strategy of nucleic acid probe assays" Elsevier, New York. Generally, highly stringent hybridization and wash conditions are selected to be about 5°C lower than the thermal melting point (T_m) for the specific sequence at a defined ionic strength and pH. Typically, under "stringent conditions" a probe will hybridize to its target subsequence, but to no other sequences.

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The T_m is the temperature (under defined ionic strength and pH) at which 50% of the target sequence hybridizes to a perfectly matched probe. Very stringent conditions are selected to be equal to the T_m for a particular probe. An example of stringent hybridization conditions for hybridization of complementary nucleic acids which have more than 100 complementary residues on a filter in a Southern or northern blot is 50% formamide with 1 mg of heparin at 42°C, with the hybridization being carried out overnight. An example of highly stringent wash conditions is 0.1 5M NaCl at 72°C for about 15 minutes. An example of stringent wash conditions is a 0.2x SSC wash at 65°C for 15 minutes (see, Sambrook, infra, for a description of SSC buffer). Often, a high stringency wash is preceded by a low stringency wash to remove background probe signal. An example medium stringency wash for a duplex of, e.g., more than 100 nucleotides, is 1x SSC at 45°C for 15 minutes. An example low stringency wash for a duplex of, e.g., more than 100 nucleotides, is 4-6x SSC at 40°C for 15 minutes. For short probes (e.g., about 10 to 50 nucleotides), stringent conditions typically involve salt concentrations of less than about 1.0 M Na ion, typically about 0.01 to 1.0 M Na ion concentration (or other salts) at pH 7.0 to 8.3, and the temperature is typically at least about 30°C. Stringent conditions can also be achieved with the addition of destabilizing agents such as formamide. In general, a signal to noise ratio of 2x (or higher) than that observed for an unrelated probe in the particular hybridization assay indicates detection of a specific hybridization. Nucleic acids that do not hybridize to each other under stringent conditions are still substantially identical if the proteins that they encode are substantially

identical. This occurs, e.g., when a copy of a nucleic acid is created using the maximum codon degeneracy permitted by the genetic code.

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The following are examples of sets of hybridization/wash conditions that may be used to clone nucleotide sequences that are homologues of reference nucleotide sequences of the present invention: a reference nucleotide sequence preferably hybridizes to the reference nucleotide sequence in 7% sodium dodecyl sulfate (SDS), 0.5 M NaPO₄, 1 mM EDTA at 50°C with washing in 2X SSC, 0.1% SDS at 50°C, more desirably in 7% sodium dodecyl sulfate (SDS), 0.5 M NaPO₄, 1 mM EDTA at 50°C with washing in 1X SSC, 0.1% SDS at 50°C, more desirably still in 7% sodium dodecyl sulfate (SDS), 0.5 M NaPO₄, 1 mM EDTA at 50°C with washing in 0.5X SSC, 0.1% SDS at 50°C, preferably in 7% sodium dodecyl sulfate (SDS), 0.5 M NaPO₄, 1 mM EDTA at 50°C with washing in 0.1X SSC, 0.1% SDS at 50°C, more preferably in 7% sodium dodecyl sulfate (SDS), 0.5 M NaPO₄, 1 mM EDTA at 50°C with washing in 0.1X SSC, 0.1% SDS at 50°C.

A "subsequence" refers to a sequence of nucleic acids or amino acids that comprise a part of a longer sequence of nucleic acids or amino acids (e.g., protein) respectively.

Substrate: a substrate is the molecule that an enzyme naturally recognizes and converts to a product in the biochemical pathway in which the enzyme naturally carries out its function, or is a modified version of the molecule, which is also recognized by the enzyme and is converted by the enzyme to a product in an enzymatic reaction similar to the naturally-occurring reaction.

Transformation: a process for introducing heterologous DNA into a plant cell, plant tissue, or plant. Transformed plant cells, plant tissue, or plants are understood to encompass not only the end product of a transformation process, but also transgenic progeny thereof.

"Transformed," "transgenic," and "recombinant" refer to a host organism such as a bacterium or a plant into which a heterologous nucleic acid molecule has been introduced. The nucleic acid molecule can be stably integrated into the genome of the host or the nucleic acid molecule can also be present as an extrachromosomal molecule. Such an extrachromosomal molecule can be auto-replicating. Transformed cells, tissues, or plants are understood to encompass not only the end product of a transformation process, but also transgenic progeny thereof. A "non-transformed," "non-transgenic," or "non-recombinant" host refers to a wild-type organism, e.g., a bacterium or plant, which does not contain the heterologous nucleic acid molecule.

Viability: "viability" as used herein refers to a fitness parameter of a plant. Plants are assayed for their homozygous performance of plant development, indicating which proteins are essential for plant growth.

I. Identification of Essential Arabidopsis thaliana Nucleotide Sequences and Encoded Proteins Using Ac/Ds Transposon or T-DNA-Mediated Mutagenesis

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As shown in the examples below, the essentiality of the nucleotide sequences described herein for normal plant growth and development, have been demonstrated for the first time in *Arabidopsis* using *Ac/Ds* transposon or T-DNA-mediated mutagenesis. Having established the essentiality of the function of the encoded proteins in *Arabidopsis thaliana* and having identified the nucleotide sequences encoding these essential proteins, the inventors thereby provide an important and sought after tool for new herbicide development.

Arabidopsis insertional mutant lines segregating for seedling lethal mutations are identified as a first step in the identification of essential proteins. Starting with T2 seeds collected from single T1 plants containing T-DNA insertions in their genomes, those lines segregating homozygous seedling lethal seedlings are identified. Ds transposon insertion lines are produced as described in Sundaresan et al. (1995) (Genes and Dev., 9:1797-1810), incorporated herein by reference. Starting with F3 or F4 seeds collected from single F2 or F3 kanamycin-resistant plants containing Ds insertions in their genomes (see Figure 3 of Sundaresan et al. (1995) (Genes and Dev., 9:1797-1810), those lines segregating homozygous seedling lethal seedlings are identified. These lines are found by placing seeds onto minimal plant growth media, which contains the fungicides benomyl and maxim, and screening for inviable seedlings after 7 and 14 days in the light at room temperature. Inviable phenotypes include altered pigmentation or altered morphology. These phenotypes are observed either on plates directly or in soil following transplantation of seedlings.

Essential genes are also identified through the isolation of lethal mutants blocked in early development. Examples of lethal mutants include those blocked in the formation of the male or female gametes or embryo. Gametophytic mutants are found by examining T1 insertion lines for the presence of 50% aborted pollen grains or ovules. Embryo defective mutants produce 25% defective seeds following self-pollination of T1 plants (see Errampalli et al. 1991, Plant Cell 3:149-157; Castle et al. 1993, Mol Gen Genet 241:504-514).

When a line is identified as segregating a seedling lethal or an embryo defective phenotype, it is determined if the resistance marker in the *Ds* transposon or T-DNA insertion co-segregates with the lethality (Errampalli *et al.* (1991) The Plant Cell, 3:149-157). Cosegregation analysis is done by placing the seeds on media containing the selective agent and scoring the seedlings for resistance or sensitivity to the agent. Examples of selective agents used are kanamycin, hygromycin, or phosphinothricin. About 35 resistant seedlings are transplanted to soil and their progeny are examined for the segregation of the seedling lethal. In the case in which the *Ds* transposon or T-DNA insertion disrupts an essential gene, there is co-segregation of the resistance phenotype and the seedling lethal or embryo defective phenotype in every plant. Therefore, in such a case, all resistant plants segregate a seedling lethal or embryo defective phenotype in the next generation; this result indicates that each of the resistant plants is heterozygous for the mutation and hemizygous for the T-DNA insert causing the mutation.

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For the *Arabidopsis* lines showing co-segregation of the transposon-encoded or T-DNA-encoded resistance marker and the lethal phenotype, PCR-based molecular approaches such as, TAIL-PCR (Liu *et al.* (1995) Plant J., 8:457-463; Liu and Whittier (1995), Genomics, 25:674-681), TAIL2k, vectorette PCR (Riley *et al.* (1990) Nucleic Acids Research, 18: 2887-2890), or the GenomeWalkerTM kit (CLONTECH Laboratories, Inc., Palo Alto, CA), may be used to directly amplify the plant DNA fragments flanking the transposon or T-DNA. Each of these techniques utilizes the known sequence of the transposon or T-DNA, and can be used to recover small (less than 5 kb) fragments directly adjacent to the insertion. PCR products are isolated and their DNA sequence is determined.

Alternatively, plasmid rescue may be used to isolate the plant DNA/T-DNA border fragments. Southern blot analysis may be performed as an initial step in the characterization of the molecular nature of each insertion. Southern blots are done with genomic DNA isolated from heterozygotes and using probes capable of hybridizing with the T-DNA vector DNA. Using the results of the Southern analysis, appropriate restriction enzymes are chosen to perform plasmid rescue in order to molecularly clone *Arabidopsis thaliana* genomic DNA flanking one or both sides of the T-DNA insertion. Plasmids obtained in this manner are analyzed by restriction enzyme digestion to sort the plasmids into classes based on their digestion pattern. For each class of plasmid clone, the DNA sequence is determined.

The resulting sequences, obtained by any of the above outlined approaches, are analyzed for the presence of non-Ds transposon and non-T-DNA vector sequences, as appropriate. When such sequences are found, they are used to search DNA and protein databases using the BLAST and BLAST2 programs (Altschul et al. (1990) J Mol. Biol. 215: 403-410; Altschul et al. (1997) Nucleic Acid Res. 25:3389-3402, both incorporated herein by reference). Additional genomic and cDNA sequences for each gene are identified by standard molecular biology procedures.

II. Recombinant Production Of Essential Proteins And Uses Thereof

For recombinant production of a protein of the invention in a host organism, a nucleotide sequence encoding the protein is inserted into an expression cassette designed for the chosen host and introduced into the host where it is recombinantly produced. The choice of the specific regulatory sequences such as promoter, signal sequence, 5' and 3' untranslated sequence, and enhancer appropriate for the chosen host is within the level of the skill of the routineer in the art. The resultant molecule, containing the individual elements linking in the proper reading frame, is inserted into a vector capable of being transformed into the host cell. Suitable expression vectors and methods for recombinant production of proteins are well known for host organisms such as *E. coli*, yeast, and insect cells (see, *e.g.*, Lucknow and Summers, *Bio/Technol.* 6:47 (1988)). Additional suitable expression vectors are baculovirus expression vectors, *e.g.*, those derived from the genome of *Autographica californica* nuclear polyhedrosis virus (AcMNPV). A preferred baculovirus/insect system is PVL1392(3) used to transfect *Spodoptera frugiperda* SF9 cells (ATCC) in the presence of linear *Autographica californica* baculovirus DNA (Phramingen, San Diego, CA). The resulting virus is used to infect HighFive *Tricoplusia ni* cells (Invitrogen, La Jolla, CA).

Recombinantly produced proteins are isolated and purified using a variety of standard techniques. The actual techniques used vary depending upon the host organism used, whether the protein is designed for secretion, and other such factors. Such techniques are well known to the skilled artisan (*see*, *e.g.* chapter 16 of Ausubel, F. *et al.*, "Current Protocols in Molecular Biology", pub. by John Wiley & Sons, Inc. (1994).

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III. Assays For Characterizing The Essential Proteins

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The recombinantly produced proteins described herein are useful for a variety of purposes. For example, they can be used in *in vitro* assays to screen known herbicidal chemicals whose target has not been identified to determine if they inhibit protein activity. Such *in vitro* assays may also be used as more general screens to identify chemicals that inhibit such protein activity and that are therefore novel herbicide candidates. Recombinantly produced proteins may also be used to elucidate the complex structure of these molecules and to further characterize their association with known inhibitors in order to rationally design new inhibitory herbicides. Alternatively, the recombinant protein can be used to isolate antibodies or peptides that modulate the activity and are useful in transgenic solutions.

IV. In vitro Inhibitor Assay: Discovery of Small Molecule Ligands That Interact with Essential Proteins Of Unknown Biochemical Function

Once a protein has been identified as a potential herbicide target based on its essentiality for normal plant growth and viability, a next step is to develop an assay that allows screening large number of chemicals to determine which ones interact with the protein. Although it is straightforward to develop assays for proteins of known function, developing assays with proteins of unknown functions can be more difficult.

To address this issue, novel technologies are used that can detect interactions between a protein and a compound without knowing the biological function of the protein. A short description of three methods is presented, including fluorescence correlation spectroscopy, surface-enhanced laser desorption/ionization, and biacore technologies.

Fluorescence Correlation Spectroscopy (FCS) theory was developed in 1972 but it is only in recent years that the technology to perform FCS became available (Madge et al. (1972) Phys. Rev. Lett., 29: 705-708; Maiti et al. (1997) Proc. Natl. Acad. Sci. USA, 94: 11753-11757). FCS measures the average diffusion rate of a fluorescent molecule within a small sample volume. The sample size can be as low as 10³ fluorescent molecules and the sample volume as low as the cytoplasm of a single bacterium. The diffusion rate is a function of the mass of the molecule and decreases as the mass increases. FCS can therefore be applied to protein-ligand interaction analysis by measuring the change in mass and therefore in diffusion rate of a molecule upon binding. In a typical experiment, the target to be analyzed is expressed as a recombinant protein with a sequence tag, such as a poly-histidine

sequence, inserted at the N or C-terminus. The expression takes place in *E. coli*, yeast or insect cells. The protein is purified by chromatography. For example, the poly-histidine tag can be used to bind the expressed protein to a metal chelate column such as Ni2+ chelated on iminodiacetic acid agarose. The protein is then labeled with a fluorescent tag such as carboxytetramethylrhodamine or BODIPY® (Molecular Probes, Eugene, OR). The protein is then exposed in solution to the potential ligand, and its diffusion rate is determined by FCS using instrumentation available from Carl Zeiss, Inc. (Thornwood, NY). Ligand binding is determined by changes in the diffusion rate of the protein.

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Surface-Enhanced Laser Desorption/Ionization (SELDI) was invented by Hutchens and Yip during the late 1980's (Hutchens and Yip (1993) Rapid Commun. Mass Spectrom. 7: 576-580). When coupled to a time-of-flight mass spectrometer (TOF), SELDI provides a mean to rapidly analyze molecules retained on a chip. It can be applied to ligand-protein interaction analysis by covalently binding the target protein on the chip and analyze by MS the small molecules that bind to this protein (Worrall et al. (1998) Anal. Biochem. 70: 750-756). In a typical experiment, the target to be analyzed is expressed as described for FCS. The purified protein is then used in the assay without further preparation. It is bound to the SELDI chip either by utilizing the poly-histidine tag or by other interaction such as ion exchange or hydrophobic interaction. The chip thus prepared is then exposed to the potential ligand via, for example, a delivery system capable to pipette the ligands in a sequential manner (autosampler). The chip is then submitted to washes of increasing stringency, for example a series of washes with buffer solutions containing an increasing ionic strength. After each wash, the bound material is analyzed by submitting the chip to SELDI-TOF. Ligands that specifically bind the target will be identified by the stringency of the wash needed to elute them.

Biacore relies on changes in the refractive index at the surface layer upon binding of a ligand to a protein immobilized on the layer. In this system, a collection of small ligands is injected sequentially in a 2-5 microlitre cell with the immobilized protein. Binding is detected by surface plasmon resonance (SPR) by recording laser light refracting from the surface. In general, the refractive index change for a given change of mass concentration at the surface layer, is practically the same for all proteins and peptides, allowing a single method to be applicable for any protein (Liedberg *et al.* (1983) Sensors Actuators 4: 299-304; Malmquist (1993) Nature, 361: 186-187). In a typical experiment, the target to be analyzed is expressed

as described for FCS. The purified protein is then used in the assay without further preparation. It is bound to the Biacore chip either by utilizing the poly-histidine tag or by other interaction such as ion exchange or hydrophobic interaction. The chip thus prepared is then exposed to the potential ligand via the delivery system incorporated in the instruments sold by Biacore (Uppsala, Sweden) to pipette the ligands in a sequential manner (autosampler). The SPR signal on the chip is recorded and changes in the refractive index indicate an interaction between the immobilized target and the ligand. Analysis of the signal kinetics on rate and off rate allows the discrimination between non-specific and specific interaction.

Another assay for small molecule ligands that interact with a polypeptide is an inhibitor assay. For example, such an inhibitor assay useful for identifying inhibitors of the products of essential plant nucleic acid sequences, such as the essential *Arabidopsis* proteins described herein, comprises the steps of:

- a) reacting an essential *Arabidopsis* protein described herein and a substrate thereof in the presence of a suspected inhibitor of the protein's function;
- b) comparing the rate of enzymatic activity of the protein in the presence of the suspected inhibitor to the rate of enzymatic activity under the same conditions in the absence of the suspected inhibitor; and
- c) determining whether the suspected inhibitor inhibits the essential *Arabidopsis* protein.

For example, the inhibitory effect on the activity of a hereindescribed essential Arabidopsis protein, may be determined by a reduction or complete inhibition of protein activity in the assay. Such a determination may be made by comparing, in the presence and absence of the candidate inhibitor, the amount of substrate used or intermediate or product made during the reaction.

V. Production of peptides

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Phage particles displaying diverse peptide libraries permits rapid library construction, affinity selection, amplification and selection of ligands directed against an essential protein (H.B. Lowman, *Annu. Rev. Biophys. Biomol. Struct.* 26, 401-424 (1997)). Structural analysis of these selectants can provide new information about ligand-target molecule interactions and

then in the process also provide a novel molecule that can enable the development of new herbicides based upon these peptides as leads.

VI. In Vivo Inhibitor Assay

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In one embodiment, a suspected herbicide, for example identified by *in vitro* screening, is applied to plants at various concentrations. The suspected herbicide is preferably sprayed on the plants. After application of the suspected herbicide, its effect on the plants, for example death or suppression of growth is recorded.

In another embodiment, an *in vivo* screening assay for inhibitors of the activity of a hereindescribed essential protein uses transgenic plants, plant tissue, plant seeds or plant cells capable of overexpressing a nucleotide sequence disclosed herein that encodes an essential protein, wherein the essential protein is enzymatically active in the transgenic plants, plant tissue, plant seeds or plant cells. A chemical is then applied to the transgenic plants, plant tissue, plant seeds or plant cells and to the isogenic non-transgenic plants, plant tissue, plant seeds or plant cells are determined after application of the chemical and compared. Compounds capable of inhibiting the growth of the non-transgenic plants, but not affecting the growth of the transgenic plants are selected as specific inhibitors of the essential protein's activity.

The invention will be further described by reference to the following detailed examples. These examples are provided for purposes of illustration only, and are not intended to be limiting unless otherwise specified.

EXAMPLES

Standard recombinant DNA and molecular cloning techniques used here are well known in the art and are described by J. Sambrook, et al., Molecular Cloning: A Laboratory Manual, 3d Ed., Cold Spring Harbor, NY: Cold Spring Harbor Laboratory Press (2001); by T.J. Silhavy, M.L. Berman, and L.W. Enquist, Experiments with Gene Fusions, Cold Spring Harbor Laboratory, Cold Spring Harbor, NY (1984) and by Ausubel, F.M. et al., Current Protocols in Molecular Biology, New York, John Wiley and Sons Inc., (1988), Reiter, et al., Methods in Arabidopsis Research, World Scientific Press (1992), and Schultz et al., Plant Molecular Biology Manual, Kluwer Academic Publishers (1998). These references describe

the standard techniques used for all steps in tagging and cloning genes from *AclDs* transposon or T-DNA mutagenized populations of *Arabidopsis*: plant infection and transformation; screening for the identification of seedling mutants; and cosegregation analysis. *Ds* transposon insertion lines produced as described in Sundaresan *et al.* (1995) Genes and Dev., 9:1797-1810) are used in these experiments. T-DNA lines are generated using vacuum infiltration or floral dip methods (Bechtold *et al.* (1993) C. R. Acad. Sci. Paris, 316:1194-1199; Clough and Bent (1998) Plant J., 16:735-743; Desfeux *et al.* (2000) Plant Physiol., 123:895-904).

Example 1: Identification of Arabidopsis Mutants with Lethal Phenotypes

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Essential genes are identified through the isolation of lethal mutants blocked in early development. Examples of lethal mutants include those blocked in the formation of the male or female gametes, embryo, or resulting seedling. Gametophytic mutants are found by examining insertion lines for the presence of 50% aborted pollen grains or ovules. Embryo defective lethal mutants usually produce 25% defective seeds following self-pollination of plants heterozygous for an insertion (see Errampalli *et al.* 1991, Plant Cell 3:149-157; Castle *et al.* 1993, Mol Gen Genet 241:504-514). Seedling lethal mutants usually segregate 25% seedlings that exhibit a lethal phenotype.

Example 2: Cosegregation Analysis for Lines with Lethal Phenotypes

The linkage of the mutation to the *Ds* or T-DNA insertion is established after identifying a transformed line segregating for a lethal phenotype of interest. A line segregating with a single functional insert will segregate for resistance in the ratio of about 2:1 (resistant: sensitive) to the selectable marker. In the case of an embryo defective mutant, one-quarter of the progeny of a plant heterozygous for an insertion will fail to germinate due to embryo lethality, resulting in a reduction of the normal 3:1 ratio to 2:1. In the case of a seedling lethal mutant, the seedlings with a mutant phenotype are excluded in the calculation of this ratio. Each of the resistant progeny is therefore heterozygous for the mutation if the *Ds* or T-DNA insertion is causing the mutant phenotype. To establish cosegregation of the insertion and the mutant phenotype, about 30 resistant progeny are transplanted to soil and each plant is shown to segregate the 25% progeny with a lethal phenotype by the appropriate screening of embryo or seedlings. When all resistant plants segregate the lethal phenotype,

there is cosegregation of the insertion and the lethal mutation and the line is designated as "tagged."

Example 3: T-DNA Border Isolation by Plasmid Rescue

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The plasmid rescue technique is used to molecularly clone Arabidopsis flanking DNA from one or both sides of the T-DNA insertion(s). Arabidopsis genomic DNA is isolated as described by Reiter et al. in Methods in Arabidopsis Research, World Scientific Press (1992). Genomic DNA is digested with a restriction endonuclease and ligated overnight. After ligation, the DNA is transformed into competent E. coli strain XL-1 Blue, DH10B, DH5 alpha, or the like, and colonies are selected on semi-solid medium containing ampicillin. Resistant colonies are picked into liquid medium with ampicillin and grown overnight. Plasmid DNA is isolated and digested with the rescue enzyme and analyzed on agarose gels containing ethidium bromide for visualization. Plasmids that represent different size classes are sequenced using primers that flank the plant DNA portion of the rescue element and the sequence is analyzed to determine what portion is plant DNA and what gene has been disrupted. The plasmid rescue is validated via PCR of template genomic DNA from a heterozygote for the insertion mutation. The experiment uses a primer anchored in the predicted flanking sequence and a primer in the T-DNA insertion. Finding a PCR product of the appropriate size, based on the sequence of the plasmid rescue clone confirms a valid rescue. Alternatively, Southern blot analysis with a probe that detects the relevant region of Arabidopsis DNA in genomic DNA from a heterozygote for the insertion mutation can be used to confirm the plasmid rescue results.

Example 4: Transposon or T-DNA Border Isolation by TAIL-PCR

Arabidopsis genomic DNA is isolated according to Reiter et al. in Methods in Arabidopsis Research, World Scientific Press (1992) or using the Nucleon PhytoPure™ Plant DNA isolation kit (Amersham International plc, Buckinghamshire, England) or the Puregene DNA isolation kit (Gentra Systems, Minneapolis, MN). Fragments of genomic DNA flanking the borders of the transposon or T-DNA are isolated using the TAIL-PCR technique (Liu et al. (1995) Plant J., 8:457-463; Liu and Whittier (1995), Genomics, 25:674-681). Three sets of 12 TAIL-PCR reactions, referred to as the primary, secondary and tertiary reactions, are performed. In each reaction, one arbitrary degenerate primer and one transposon-specific or

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T-DNA-specific primer are used. The arbitrary degenerate primer is chosen from among seven primers, LWAD1, CA50, CA51, CA52, CA53, CA54, and CA55 (Table 1), which are used to prime the genomic DNA flanking the insertion. Alternatively, less than 12 TAIL-PCR reactions are done using fewer arbitrary degenerate primers. These degenerate primers are used in combination with two sets of three, nested, transposon-specific primers (Table 2) or T-DNA-specific primers (Table 3). The transposon-specific primers are homologous to regions of the Ds elements that lie at the outermost ends of the transposons, DS5 at the 5' end (primers 5A, 5B, and 5C) and DS3 at the 3' end (primers 3A, 3B, and 3C). The T-DNAspecific primers are homologous to regions of the T-DNA that lie in the borders of the T-DNAs. For the pCSA104 and pDAP101 T-DNAs, right borders are recovered with CA66 (primary primer), CA67 (secondary primer), and CA68 (tertiary primer) and left borders are recovered with JM33 (tertiary primer); JM34 (secondary primer); and JM35 (primary primer). For the pCSA110 T-DNA, right borders are recovered with QRB1 (primary primer), QRB2 (secondary primer), and QRB3 (tertiary primer) and left borders are recovered with JM33 (tertiary primer); JM34 (secondary primer); and JM35 (primary primer). For the pPCVICEn4HPT (Hayashi et al. (1992), Science, 258:1350-1353) and pSKI015 (Weigel et al. (2000) Plant Physiol. 122:1003-1014) T-DNAs, left borders are recovered with SKI1 (primary primer), SKI2 (secondary primer), and SKI3 (tertiary primer). When the degenerate and nested primer pairs are used in a series of low and high-stringency PCR amplifications, as described in the TAIL-PCR protocol (Liu and Whittier (1995), Genomics, 25:674-681), DNA fragments are produced that correspond to the genomic DNA that is directly adjacent to the transposon or T-DNA insertion. The nucleic acid sequences of the PCR products from the tertiary TAIL-PCR reactions are then determined by standard molecular biology techniques. The resulting sequences are analyzed for the presence of non-Ds transposon or non-T-DNA vector sequence.

To confirm the integrity of the resultant products, PCR primers specific to the flanking genomic region are designed and used in conjunction with the tertiary nested primer in a PCR reaction, to confirm the transposon or T-DNA insertion point within the genomic DNA. Finding a PCR product of the appropriate size, based on the sequence of the TAIL-PCR clone confirms a valid rescue.

Table 1: Arbitrary Degenerate Primers

	SEQ ID NO:	<u>Primer</u>	Degen.	Primer Sequence
	101	LWAD1	1026	ngt tgw gna twt sgw gnt
	102	CA50	128	ngt cga swg ana wga a
5	103	CA51	128	tgw gna gsa nca sag a
	104	CA52	128	agw gna gwa nca wag g
	105	CA53	256	stt gnt ast nct ntg c
	106	CA54	64	ntc gas twt sgw gtt
	107	CA55	256	wgt gna gwa nca nag a

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Table 2: Nested Primers For Ds Lines

	SEQ ID NO:	<u>Primer</u>	Primer Sequence
	108	5A	actagetetacegttteegtttae
	109	5B	ttacctcgggttcgaaatcgatcgggataa
15	110	5C	aaaatcggttatacgataacggtcggtacggga
	111	3A	gggtcttgcggatctgaatatatgttttcatgtgtg
	112	3B	taccgaagaaaaataccggttcccgtccgatttcgac
	113	3C	ggatcgtatcggttttcgattaccgtatttatcc

20 Table 3: Nested Primers For T-DNA Lines

	SEQ ID NO:	<u>Primer</u>	Primer Sequence
	114	CA66	att agg cac ccc agg ctt tac act tta tg
	115	CA67	gta tgt tgt gtg gaa ttg tga gcg gat aac
	116	CA68	taa caa ttt cac aca gga aac agc tat gac
25	117	JM33	tag cat ctg aat ttc ata acc aat ctc gat aca c
	118	JM34	get tee tat tat ate tte cea aat tae eaa tae a
	119	JM35	gcc ttt tca gaa atg gat aaa tag cct tgc ttc c
	120	QRB1	caa act agg ata aat tat cgc gcg cgg tgt ca
	121	QRB2	ggt gtc atc tat gtt act aga tcg gga att ga
30	122	QRB3	cgc cat ggc ata tgc tag cat gca taa ttc
	123	SKI1	aat tgg taa tta ctc ttt ctt ttc ctc cat att ga
	124	SKI2	ata ttg acc atc ata ctc att gct gat cca t
	125	SKI3	tga tcc atg tag att tcc cgg aca tga a

Example 5: Transposon or T-DNA Border Isolation by TAIL2k PCR

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Arabidopsis genomic DNA is isolated according to Reiter et al. in Methods in Arabidopsis Research, World Scientific Press (1992) or using the Nucleon PhytoPure™ Plant DNA isolation kit (Amersham International plc, Buckinghamshire, England) or the Puregene DNA isolation kit (Gentra Systems, Minneapolis, MN). Fragments of genomic DNA flanking the borders of the transposon or T-DNA are isolated using the TAIL2k PCR technique. Two sets of 12 TAIL-PCR reactions, referred to as the primary and secondary reactions, are performed. In each reaction, one arbitrary degenerate primer and one transposon-specific or T-DNA-specific primer are used. The arbitrary degenerate primer is selected from among six primers; CA50, CA51, CA52, CA53, CA54, and CA55 (Table 1), which are used to prime the genomic DNA flanking the insertion. Alternatively, less than 12 TAIL-PCR reactions are done using fewer arbitrary degenerate primers. These degenerate primers are used in combination with two sets of two, nested, transposon-specific primers (Table 2) or T-DNAspecific primers (Table 3). The transposon-specific primers are homologous to regions of the Ds elements that lie at the outermost ends of the transposons, DS5 at the 5' end (primers 5A). 5B, and 5C) and DS3 at the 3' end (primers 3A, 3B, and 3C). The T-DNA-specific primers are homologous to regions of the T-DNA that lie in the borders of the T-DNAs. For the pCSA104 and pDAP101 T-DNAs, right borders are recovered with CA66 (primary primer). CA67 (secondary primer), and CA68 (sequencing primer) and left borders are recovered with JM33 (sequencing primer), JM34 (secondary primer), and JM35 (primary primer). Primers CA66, CA67, and CA68 are also known as RB1, RB2, and RB3, respectively. Primers JM35. JM34, and JM33 are also known as LB1, LB2, and LB3, respectively. For the pCSA110 T-DNA, right borders are recovered with QRB1 (primary primer), QRB2 (secondary primer), and QRB3 (sequencing primer) and left borders are recovered with JM33 (sequencing primer); JM34 (secondary primer); and JM35 (primary primer). For the pPCVICEn4HPT (Hayashi et al. (1992), Science, 258:1350-1353) and pSKI015 (Weigel et al. (2000) Plant Physiol. 122:1003-1014) T-DNAs, left borders are recovered with SKI1 (primary primer), SKI2 (secondary primer), and SKI3 (sequencing primer). When the degenerate and nested primer pairs are used in a series of low and high-stringency PCR amplifications, as described in the TAIL-PCR protocol (Liu and Whittier (1995), Genomics, 25:674-681), DNA fragments are produced that correspond to the genomic DNA that is directly adjacent to the transposon

or T-DNA insertion. TAIL2k-PCR differs from the original TAIL-PCR protocol by the elimination of the tertiary PCR and modification of the secondary PCR. The cycling conditions used in the secondary reaction are modified to include 5 high annealing temperature cycles (64 degrees C) at the beginning, three additional so-called super cycles, and five additional low annealing temperature cycles (44 degrees C) at the end of the reaction. The melting and extension times are the same as all other TAIL-PCR reactions. Additionally, the reaction volume is increased to 40 microliters. The nucleic acid sequences of the PCR products from the secondary TAIL2k-PCR reactions are then determined by standard molecular biology techniques. The resulting sequences are analyzed for the presence of non-Ds transposon or non-T-DNA vector sequence.

To confirm the integrity of the resultant products, PCR primers specific to the flanking genomic region are designed and used in conjunction with the tertiary nested primer in a PCR reaction, to confirm the transposon or T-DNA insertion point within the genomic DNA. Finding a PCR product of the appropriate size, based on the sequence of the TAIL2k-PCR sequencing result confirms a valid rescue.

Example 6: Identification of Both Borders of a T-DNA or Ds Insertion

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If the results of border rescue provide information on only one of the two borders for an insertion in a given line, additional experiments are performed to identify the second border. These experiments are necessary to show that a single gene has been disrupted in a given line. In some cases, an insertion can affect more than a single gene due to a chromosomal deletion or rearrangement. In those cases, additional experiments are required to identify which of the affected genes is responsible for the lethal phenotype.

When both borders of an insertion are not recovered, primers are designed to isolate a PCR product that will provide information on the location of the missing border. Three primers are chosen in *Arabidopsis* genomic DNA on the opposite side of the insertion about one, two, and five kb away from the insertion point; the primers point towards the expected second border. Long PCR conditions (Advantage 2, Clontech) are then employed following the manufacturer's directions to amplify the relevant region from genomic DNA isolated from a heterozygote for the lethal mutation. PCR reactions are performed using appropriate pairs of genomic and T-DNA or *Ds* border primers. Finding a PCR product of the appropriate size.

based on the sequence of the TAIL-PCR clone confirms a valid rescue of the second border. In some cases, the PCR product is directly sequenced to determine the exact insertion point.

If the second border is not recovered with this method, an additional set of PCR reactions are preformed. In these experiments, the genomic primers are paired with a series of internal T-DNA or *Ds* primers designed at about one kb intervals in both orientations across the entire T-DNA or *Ds* vector sequence. Finding a PCR product of the appropriate size, based on the sequence of the TAIL-PCR clone confirms a valid rescue of the second border. In some cases, the PCR product is directly sequenced to determine the exact insertion point. Any borders recovered with this approach are classified as abnormal because they lack the ends of the *Ds* transposon or the expected 24 bp T-DNA imperfect repeat characteristic of right and left borders.

Example 7: Identification of Insertion Points for Lines with Lethal Phenotypes

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For each line with a lethal phenotype, the sequences of the borders of the insertion are determined and the insertion points in the *Arabidopsis* genome are deduced. For *Ds* insertion lines, PCR products are obtained from the Ds3 and Ds5 borders. For T-DNA lines, PCR products or plasmid rescue clones are obtained from left (LB), right (RB), or abnormal (AB) borders. These sequences are used in BLASTn searches against nucleotide databases (Altschul *et al.* (1990) J Mol. Biol. 215:403-410; Altschul *et al.* (1997) Nucleic Acids Res. 25:3389-3402). The results are summarized in Table 4. *Ds* line names begin with ET or GT; T-DNA line names are numbers. The insertion point (Insert Pt.) and the direction of the flanking sequence (Dir.) either up (U) or down (D) in the genome section is noted. Often, small deletions or duplications of genomic DNA accompany the insertion of a T-DNA or *Ds* transposon.

The gene that has been inactivated in a given line with a lethal phenotype is determined from the insertion points for that line. Often, the precise location of an ORF for a given gene is not known, but predictions are available in genome sections deposited in GenBank. The precise boundaries of that ORF is determined as described in Example 7.

Table 4: Insertion Points For Lines With Lethal Phenotypes

Gene	Line#	Border	Genome Section	Acc. #	Insert Pt.	Dir.
942	942	LB	K24G6	AB012242	33667	D

978	978	LB	F23N20	AC016972	58221	D
	978	LB	F23N20	AC016972	58301	U
3218	3218	LB	T8K14	AC007202	10500	D
	3218	LB	T8K14	AC007202	10540	U
4563	4563	LB	ATCHRII092	AC006438	25542	D
8794	8794	LB	F2J6	AC009526	45854	$\frac{\tilde{D}}{D}$
	8794	LB	F2J6	AC009526	45879	Ū
9106	9106	LB	T2J13	AL132967	78013	Ū
	9106	AB	T2J13	AL132967	77943	$\frac{0}{D}$
10708	10708	RB	F1I21	AC005687	40005	D
10700	10708	LB	F1I21	AC005687	40042	U
	70241	LB	F1I21	AC005687	40210	D
	70241	RB	F1I21	AC005687	40215	U
10844	10844	LB	F13F21	AC007504	60873	U
10044	10844	LB	F13F21	AC007504	60839	D
10951	10951	LB	MKP11	AB005238	20298	D
10931	10951	LB	MKP11	AB005238		<u>U</u>
12935		LB			20318	
12933	12935 12935	LB LB	ATCHRII150 ATCHRII150	AC005168	36510	D
12002			T27G7	AC005168	36545	U
13823	11361	LB		AC006932	78096	<u>n</u>
	11361	AB	T27G7	AC006932	78065	D
	13823	LB	T27G7	AC006932	78096	<u>U</u>
14510	13823	RB	T27G7	AC006932	77722	<u>D</u>
14519	14519	LB	ATCHRIV72	AL161576	50259	<u>U</u>
14610.1	14519	AB	ATCHRIV72	AL161576	50228	D
14610.1	14610.1	LB	F4P13	AC009325	55319	U
	14610.1	RB	F4P13	AC009325	55442	D
14891	14891	<u>LB</u>	ATCHRIV89	AL161593	11412	U
	14891	RB	ATCHRIV89	AL161593	11313	<u>D</u>
14986	14986	LB	K10D20	AP000410	51816	D
	14986	RB	K10D20	AP000410	54505	U
15377	15377	RB	F28G11	AC074025	19572	D
	15377	LB	F28G11	AC074025	19587	U
16219	16219	LB	MRO11	AB005244	51998	U
	16219	LB	MRO11	AB005244	51995	D
16547	16547	LB	ATCHRIV65	AL161565	80692	D
	16547	RB	ATCHRIV65	AL161565	80791	U
20933	20933	LB	ATCHRII146	AC004747	47678	D
	20933	LB	ATCHRII146	AC004747	47683	U
21455	21455	LB	ATCHRIV54	AL161554	105596	U
	21455	RB	ATCHRIV54	AL161554	105542	D
21878	21878	LB	T19F11	AC009918	19609	D
23915	23915	LB	ATCHRII008	AC005936	49629	D
	23915	LB	ATCHRII008	AC005936	49657	Ū
30945	30945	LB	ATCHRII192	AC004238	2411	D
	30945	LB	ATCHRII192	AC004238	2410	U
				110001200		

31895	31895	LB	MTI20	AB013396	52020	D
	31895	LB	MTI20	AB013396	52089	Ū
34269	34269	LB	T4O12	AC007396	92811	Ū
	34269	RB	T4O12	AC007396	92808	D
34540	34540	LB	T1G11	AC002376	41572	D
	34540	LB	TIG11	AC002376	41608	Ū
	72902	LB	T1G11	AC002376	41494	U
	72902	LB	T1G11	AC002376	41465	D
34555	34555	LB	T1F15	AC004393	42152	D
	54334	RB	T1F15	AC004393	41803	U
	54334	LB	T1F15	AC004393	41671	D
35154	35154	RB	MWD9	AB007651	45718	D
55151	35154	LB	MWD9	AB007651	45732	Ū
35438	35438	LB	MAL21	AP000383	25170	D
JJ+J0	35438	LB	MAL21	AP000383	25738	U
37351	37351	LB	F25C20	AC007296	52890	U
37331	37351	RB	F25C20	AC007296	52196	D
37389	37389	LB	F3F19	AC007357	45488	U
31307	37389	RB	F3F19	AC007357	45471	D
38108	·38108	LB	ATCHRII150	AC007357 AC005168	83430	<u>Б</u>
36106	38108	RB	ATCHRII150	AC005168 AC005168	83446	<u>น</u>
43301	43301	RB	T22D16	AL357612	57549	D
43301	43301	LB	T22D16	AL357612 AL357612	57599	<u>บ</u>
46250	46250	LB	F17A9	AC016827	74222	
40230	46250	RB	F17A9	AC016827 AC016827	74274	D U
47050A	47050	LB	T23E18	AC010827 AC009978	49445	D
47030A	47050	RB	T23E18	AC009978 AC009978		<u>น</u>
52949A	52949	LB	K16H17	AB016884	49475	
J4747A	52949	LB LB	K16H17	AB016884	34713 34718	D U
53210A	53210	RB	ATCHRII017	AC007167		
JJZIUA	53210	LB	ATCHRII017		92796	D U
	69121	LB	··	AC007167	92942	
	69121	LB	ATCHRII017	AC007167	94478	D
55483	55483	RB	ATCHRII017 ATCHRII164	AC007167	94502	U
<u> </u>				AC005727	71269	<u>U</u>
50251 A	55483	LB	ATCHRII164	AC005727	71258	<u>D</u>
58351A	58351	RB	MYH9	AB016893	42547	D
(0044	58351	LB	MYH9	AB016893	42772	U
60944	60944	LB	F1B16	AC023754	89492	U
60007	60944	LB	F1B16	AC023754	89428	D
62837	62837	<u>LB</u>	T21J18	AL132963	70906	<u>U</u>
(5210	62837	LB	T21J18	AL132963	70873	<u>D</u>
65310	65310	<u>LB</u>	T20H2	AC022472	8158	U
(0101	65310	<u>RB</u>	T20H2	AC022472	8096	D
68181	68181	RB	F12A12	AL133314	38270	U
	68181	<u>LB</u>	F12A12	AL133314	38275	<u>D</u>
70913	70913	LB	T24H18	AL353013	5347	D

	70913	LB	T24H18	AL353013	5358	U
71067	71067	LB	F2E2	AC069252	63031	U
	71067	LB	F2E2	AC069252	62932	D
71654	71654	RB	MYA6	AB023046	71956	U
	71654	LB	MYA6	AB023046	71907	D
ET3172	ET3172	DS5	ATCHRIV4	AL161492	134442	U
ET3546	ET3546	DS3	ATCHRII115	AC006081	20874	D
	ET3546	DS5	ATCHRII115	AC006081	20973	U

Example 8: Identification of cDNAs for Essential Genes

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A cDNA for a gene identified as essential is identified using a variety of approaches. This information enables the ORF for a given gene to be identified and used for other experiments including expression of the corresponding protein in heterologous systems.

If there is a full-length cDNA deposited in GenBank or published elsewhere, that sequence may be checked independently using methods described below. Alternatively, the sequence may be considered to be correct.

In some cases, there are published EST sequences that can be assembled to cover the entire ORF from start codon to stop codon. This sequence may be checked independently using methods described below or it may be considered to be correct.

Often part of the cDNA is published and this information can be used to identify the entire ORF. If the 5' end containing the start codon is known, 3' RACE is performed to identify the remainder of the cDNA. If the 3' end containing the stop codon is known, 5' RACE is performed to identify the remainder of the cDNA. If both the 5' and the 3' ends are known, but the sequence between the two ends of the cDNA is not known, PCR is performed with primers hybridizing to each end of the cDNA. In all three of these cases, PCR is performed using template DNA from a GeneRacer (Invitrogen) or a Marathon (Clontech) cDNA library prepared from RNA isolated from seedling tissue. A resulting PCR product is TA-cloned (Original TA-Cloning kit, Invitrogen) and sequenced.

If no part of the cDNA is published, the cDNA is identified by starting from gene model predictions in the annotation for genomic clones or elsewhere. To identify the ORF, primers are designed to the 5' and 3' ends of the predicted ORF. PCR is performed using template DNA from a cDNA library prepared from seedling tissue or the pFL61 *Arabidopsis* cDNA library (Minet *et al.* (1992) Plant J. 2: 417-422). The resulting PCR product is TAcloned (Original TA-Cloning kit, Invitrogen) and sequenced. Alternatively, 5' and 3' RACE are performed with primers predicted by gene models to be in exons. PCR is performed using

template DNA from a GeneRacer (Invitrogen) or a Marathon (Clontech) cDNA library prepared from RNA isolated from seedling tissue. A resulting PCR product is TA-cloned (Original TA-Cloning kit, Invitrogen) and sequenced.

If the cDNA sequence is the same as the sequence predicted in the GenBank annotation, the experiments confirm for the first time the actual ORF. If the cDNA sequence is not the same as the sequence predicted in the GenBank annotation, the experiments identify for the first time the actual ORF. In some cases, more than one cDNA sequence is found for a given gene and both sequences are included in this application.

10 Example 9: Description of Essential Genes

The putative function of the protein encoded by each essential gene is determined from analysis of the ORF in each cDNA. Information from the relevant *Arabidopsis* genomic section deposited in GenBank is used as a starting point to explore the function of a given gene. This analysis also includes BLAST searches (Altschul *et al.* (1990) J. Mol. Biol. 215:403-410; Altschul *et al.* (1997) Nucleic Acids Res. 25:3389-3402) of sequence databases to identify similar proteins. Table 5 describes the putative functions for the essential genes discovered in this application.

Table 5: Putative Functions For Essential Genes

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Gene	SEQ ID Nos:	Putative Function & Similar Genes	References
00942	1-2	similarity to disease resistance protein large gene family in Arabidopsis including disease resistance proteins RPP1-WsA,B&C similar to tobacco TMV resistance protein N	Whitham, S. et al. (1994) Cell 78:1101-1115; Botella, M.A., et al. (1998) Plant Cell 10: 1847-1860
00978	3-4	unknown protein similar to <i>Arabidopsis</i> protein of unknown function (CAB87660) & ESTs from many plants	none
03218	5-6	AAA ATPase similar to <i>E. coli</i> FtsH cell division protein (P28691) that acts as an ATP-dependent metallopeptidase; homologs in many species	Schumann, W. (1999) FEMS Microbiol Rev 23:1-11; Langer, T. (2000) Trends Biochem Sci 2000 25:247-251

04563	7-8	unknown protein large gene family in Arabidopsis	none
		of unknown function proteins	
08794	9-10	putative histidine decarboxylase similar to Brassica, tomato (tom92), and rice putative histidine decarboxylases	Picton, S et al. (1993) Plant Mol Biol 23:627-631; Watanabe, T et al. (1990) Trends Pharmacol Sci 11:363- 367; Vaaler, G.L. & Snell, E.E. (1989) Biochemistry 28:7306-7313
09106	11-12	cytosolic 40S ribosomal protein S11-alpha	Browning, K.S. (1996) <i>Plant Mol Biol</i> 32:107-144; Gantt, J. S. & Thompson, M.D. (1990) <i>J. Biol Chem</i> 265:2763-2767
10708	13-14	cytoplasmic 60S ribosomal protein L3	Peltz, S.W. et al. (1999) Mol Cell Biol 19:384-391; Kim, Y. et al. (1990) Gene 93:177- 182; Wickner, R.B et al. (1982) Proc Natl Acad Sci USA 79:4706-4708
10844	15-16	40S ribosomal protein S17-like	Gantt, J.S. & Thompson, M.D. (1990) J Biol Chem 265:2763-2767; Wiener, L. et al. (1988) Nucleic Acids Res 16:1233-1250
10951	17-18	phytoene synthase	Welsch, R. et al. (2000) Planta 211:846-854; Shewmaker, C.K. et al. (1999) Plant J. 20:401-412; Von Lintig, J. et al. (1997) Plant J. 12:625-634
12935	19-20	putative choline kinase similar to soybean choline kinase (T08815) and mouse & human choline/ethanolamine kinases	Monks, D.E. et al. (1996) Plant Physiol. 110:1197-1205; Bligny, R. et al. (1989) J Biol Chem. 264:4888-4895; Wharfe, J. & Harwood, J.L. (1979) Biochim Biophys Acta. 575:102-111
13823	21-22	magnesium protoporphyrin IX chelatase subunit D	Papenbrock, J. et al. (1997) Plant J. 12:981-990; Papenbrock, J. et al. (2000) Plant Physiol. 122:1161-1169; Luo, M. et al. (1999) Plant Mol Biol. 41:721-731; Jensen, P.E. et al. (1996) Mol. Gen. Genet. 250:383-394

14519	23-24	putative protein	none
1.517	23 2.	small gene family in Arabidopsis	none
		of unknown function proteins	
14610.1	25-26	putative cell division control protein; similar to cdc48, AAA ATPase proteins similar to S. pombe AAA ATPase (CAB16902); Arabidopsis cdc48 homolog (P54609); cdc48/valosin-containing protein homologs from soybean, Capsicum annuum, rice, Dictylostelium; Drosophila smallminded	Frohlich, K.U. et al. (1991) J Cell Biol. 114:443-453; Feiler, H.S. et al. (1995) EMBO J. 14:5626-5637; Langer, T. (2000) Trends Biochem Sci 2000 25:247-251
14891	27-28	putative protein contains PFAM 02536 mTERF (mitochondrial transcription termination factor) domain; large gene family in <i>Arabidopsis</i> of unknown function proteins	Fernandez-Silva, P. et al. (1997) EMBO J 16:1066-1079
14986	29-30	ubiquitin isopeptidase T (aka ubiquitin-specific protease 14)	Wilkinson, K.D. et al. (1995) Biochemistry 34:14535- 14546; Falquet, L. et al. (1995) FEBS Lett 376:233- 237; Lindsey, D.F. et al. (1998) J Biol Chem 273:29178-29187
15377	31-32	putative formyl transferase similar to B. napus methionyl tRNA transformylase Fmt protein (AJ245479) & B. rapa S-locus protein 8 (AB022076)	Cui Y et al. (1999) Plant Cell. 11:2217-2231; Suzuki, G. et al. (1999) Genetics 153:391- 400; Cusack S. (1999) Curr Opin Struct Biol. 9:66-73
16219	33-34	polyadenylation cleavage/specificity factor 100 kDa subunit (AF283277)	Bilger, A. et al. (1994) Genes Dev. 8:1106-1116; Bienroth, S. et al. (1993) EMBO J. 12:585-594; Jenny, A. et al. (1994) Mol Cell Biol. 14:8183-8190
16547	35-36	similarity to UV-induced protein Uvi31, S. pombe, G1381578 unknown function, but similar to Pectobacterium chrysanthemi SufE protein (AJ301654) involved in iron metabolism, S. pombe uvi31 protein of the BolA / YRBA family (Q12238), Synechocystis hypothetical 17.7 KDA protein SLR1419 (P74523)	Kim, S.H. et al. (1997) Environ Mol Mutagen 30:72- 81; Santos, J.M. et al. (1999) Mol Microbiol 32:789-798

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37351	59-60	strong similarity to obtusifoliol 14- alpha demethylase (CYP51; P93846) from Sorghum bicolor (also wheat & rice), member of the PF100067 cytochrome P450 family	Kushiro, M. et al. (2001) Biochem Biophys Res Commun. 285:98-104; Bak et al. (1997) Plant J. 11:191-201; Grausem, B. (1995) Plant J. 7:761-770
37389	61-62	similar to human GLE1-like required for poly(A)+ RNA export (AAC25561)	Watkins, J.L. et al. (1998) Proc. Natl. Acad. Sci. U.S.A. 95:6779-6784; Murphy, R. & Wente, S.R. (1996) Nature 383:357-360
38108	63-64	Arabidopsis 4-(cytidine 5'- phospho)-2-C-methyl-D-erythritol kinase (aka ispE & 4- diphosphocytidyl-2-C-methyl- Derythritol kinase) (AF288615) similar to E. coli ychB (aka ispE) gene (P24209)	Rohdich, F. et al. (2000) Proc Natl Acad Sci U.S.A. 97:8251-8256; Luettgen, H. et al. (2000) Proc. Natl. Acad. Sci. U.S.A. 97:1062-1067; Lange, B.M. & Croteau, R. (1999) Proc Natl Acad Sci U.S.A. 96:13714-13719
43301	65-66	similar to hypothetical bacterial proteins, including <i>Pseudomonas</i> aeruginosa protein PA0292 (F83608) & Lactococcus lactis (AAK05795)	none
46250	67-68	hypothetical protein weak similarity to hypothetical proteins from Arabidopsis (AAG51506) and mouse (BAB23375)	none
47050A	69-70	unknown protein weak similarity to Botrytis cDNA (AL115827)	none
52949A	71-72	6-phosphogluconolactonase-like protein similar to 6-phosphogluconolactonases such as human (O95336), Brassica carinata (AAK50346), & Mycobacterium tuberculosis (devB, CAB09261)	Collard, F. et al. (1999) FEBS Lett. 459:223-226; Bauer, H.P. et al. (1983) Eur J Biochem. 133:163-168
53210A	73-74	putative heat shock protein in hsp90 family similar to rye hsp82 (S65776), Ipomoea nil hsp83 (P51819), chicken hsp90 beta (Q04619) and others	Felsheim, R.F. & Das, A. (1992) Plant Physiol. 100:1764-1771; Coates, A.R. et al. (1999) Biotechnol Genet Eng Rev 16:393-405; Milioni, D. & Hatzopoulos, P. (1997) Plant Mol Biol 35:955-961

55483	75-76	putative para-aminobenzoate synthase and glutamine amidotransferase, a bifunctional enzyme similar to Streptomyces pristinaespiralis papA (AAC44866), E. coli pabB (P05041) & pabA (P00903), and Bacillus stearothermophilus anthranilate synthase component I trpE (AAD33791)	Goncharoff, P. & Nichols, B.P. (1984) J Bacteriol. 159:57-62.; Roux, B. & Walsh, C.T. (1992) Biochemistry. 31:6904-6910; Kaplan, J.B. & Nichols, B.P. (1983) J Mol Biol 168:451- 468
58351A	77-78	26S proteasome p55 protein-like similar to human 26S proteasome regulatory complex chain p55 (BAA19749), S. cerevisiae 26S proteasome regulatory complex chain RPN5 (S67695), and others	Saito, A. et al. (1997) Gene 203:241-250; Glickman, M.H. et al. (1998) Mol Cell Biol 18:3149-3162
60944	79-80	similar to Guillardia theta chloroplast 50S ribosomal protein L31 (O46917)	Yamaguchi, K. & Subramanian, A.R. (2000) <i>J Biol Chem</i> , 275:28466-28482
62837	81-82	AtClpC: regulatory subunit of Clp protease with ATPase activity (BAA82062)	Adam, Z. (2000) Biochimie 82:647-654; Sokolenko, A. et al. (1998) Planta 207:286-295; Nakabayashi, K. et al. (1999) Plant Cell Physiol. 40:504- 514; Maurizi, M.R. et al. (1990) J Biol Chem. 265:12536-12545
65310	83-84	26S proteasome regulatory subunit S3, contains a PCI PFI01399 domain similar to 26S proteasome regulatory subunit S3 from Nicotiana tabacum (P93768), carrot (Q06364), human (O43242), S. cerevisiae RPN3 (P40016), and others	Voges, D. et al. (1999) Ann Rev Biochem 68:1015-1068; Fu, H. et al. (1999) Mol Biol Rep 26:137-146; Fu, H. et al. (1999) Plant J 18:529-539; Kominami, K. et al. (1997) Mol Biol Cell 8:171-187
68181	85-86	small zinc finger-like protein TIM9 similar to mitochondrial import inner membrane translocase subunit TIM9 from several plants and S. cerevisiae (074700)	Koehler, C.M. et al. (1998) EMBO J. 17:6477-6486; Tokatlidis, K. et al. (2000) Biochem Soc Trans 28:495- 499
70913	87-88	Arabidopsis CCAAT binding protein/transcription factor Hap2a (CAA74048)	Edwards, D. et al. (1998) Plant Physiol 117:1015-1022; Albani, D. & Robert, L.S. (1995) Gene 167:209-213

71067	89-90	hypothetical protein gene family in <i>Arabidopsis</i> of unknown function proteins	none
71654	91-92	poly(A) binding protein-like	Hilson, P. et al. (1993) Plant Physiol 103:525-533; Belostotsky, D.A. & Meagher, R.B. (1993) Proc Natl Acad Sci U.S.A. 90:6686-6690; Gallie, D.R. (1998) Gene 216:1-11
ET3172	93-94	hypothetical protein small gene family in <i>Arabidopsis</i> (T47999 & T02193) of unknown function	none
ET3546	95-96	cdc27/nuc2-like protein, may contain TPR-repeat similar to human cdc27 (P30260), S. pombe nuc2 (P10505), S. cerevisiae cdc23 (P16522), and others	Hirano, T. et al. (1988) J. Cell Biol. 106:1171-1183; Chen, P.L. et al (1995) Cell Growth Differ. 6:199-210

Example 10: Expression of Recombinant Essential Proteins in E. coli

The coding region of each of the essential proteins, corresponding to cDNA clones of odd-numbered SEQ ID NO:1-96, is subcloned into an appropriate expression vector, and transformed into *E. coli* using the manufacturer's conditions. Specific examples include plasmids such as pBluescript (Stratagene, La Jolla, CA), pFLAG (International Biotechnologies, Inc., New Haven, CT), and pTrcHis (Invitrogen, La Jolla, CA). *E. coli* is cultured, and expression of the essential protein is confirmed. Recombinant protein is isolated using standard techniques.

Example 11: In Vitro Binding Assays

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Recombinant protein for each of the essential genes described in this application is obtained, for example, according to Example 10. The protein is immobilized on chips appropriate for ligand binding assays using techniques that are well known in the art. The protein immobilized on the chip is exposed to sample compound in solution according to methods well know in the art. While the sample compound is in contact with the immobilized protein, measurements capable of detecting protein-ligand interactions are conducted. Examples of such measurements are SELDI, biacore and FCS, described above. Compounds

found to bind the protein are readily discovered in this fashion and are subjected to further characterization.

The above-disclosed embodiments are illustrative. This disclosure of the invention will place one skilled in the art in possession of many variations of the invention. All such obvious and foreseeable variations are intended to be encompassed by the present invention.

CLAIMS:

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1. A method of identifying a herbicidal compound, comprising:

- a) combining a polypeptide comprising an amino acid sequence at least 90% identical to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96 with a compound to be tested for the ability to bind to said polypeptide, under conditions conducive to binding;
- b) selecting a compound identified in (a) that binds to said polypeptide;
- c) applying a compound selected in (b) to a plant to test for herbicidal activity; and
- d) selecting a compound identified in (c) that has herbicidal activity.
- 2. The method according to claim 1, wherein said polypeptide comprises an amino acid sequence at least 95% identical to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96.

3. The method according to claim 2, wherein said polypeptide comprises an amino acid sequence at least 99% identical to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96.

- 4. The method according to claim 3, wherein said polypeptide comprises an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96.
 - 5. A method of identifying a herbicidal compound, comprising:
 - c) combining a polypeptide comprising an amino acid sequence at least 90% identical to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96 with a compound to be tested for the ability to inhibit the activity of said polypeptide, under conditions conducive to inhibition;
 - d) selecting a compound identified in (a) that inhibits the activity of said polypeptide;
 - c) applying a compound selected in (b) to a plant to test for herbicidal activity; and
 - d) selecting a compound identified in (c) that has herbicidal activity.

6. The method according to claim 5, wherein said polypeptide comprises an amino acid sequence at least 95% identical to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96.

- 5 7. The method according to claim 6, wherein said polypeptide comprises an amino acid sequence at least 99% identical to an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96.
- 8. The method according to claim 7, wherein said polypeptide comprises an amino acid sequence selected from the group consisting of the even numbered SEQ ID NOs:2-96.
 - 9. A method for killing or inhibiting the growth or viability of a plant, comprising applying to the plant a herbicidal compound identified according to the method of claim 1.
- 15 10. A method for killing or inhibiting the growth or viability of a plant, comprising applying to the plant a herbicidal compound identified according to the method of claim 5.

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Arg Leu Tyr Val Thr Met Lys Glu Gly Phe Pro Leu Glu Tyr Ile Val Asp Ile Pro Leu Asp Pro Tyr Leu Phe Glu Thr Ile Cys Asn Ala Gly . 410 Val Glu Val Asp Leu Gln Lys Arg Gln Ile His Tyr Phe Met Lys Val Phe Ile Ala Leu Leu Pro Gly Ile Leu Ile Leu Trp Phe Ile Arg Glu Ser Ala Met Leu Leu Leu Ile Thr Ser Lys Arg Phe Leu Tyr Lys Lys Tyr Asn Gln Leu Phe Asp Met Ala Tyr Ala Glu Asn Phe Ile Leu Pro Val Gly Asp Val Ser Glu Thr Lys Ser Met Tyr Lys Glu Val Val Leu Gly Gly Asp Val Trp Asp Leu Leu Asp Glu Leu Met Ile Tyr Met Gly Asn Pro Met Gln Tyr Tyr Glu Lys Asp Val Ala Phe Val Arg Gly Val Leu Leu Ser Gly Pro Pro Gly Thr Gly Lys Thr Leu Phe Ala Arg Thr Leu Ala Lys Glu Ser Gly Leu Pro Phe Val Phe Ala Ser Gly Ala Glu Phe Thr Asp Ser Glu Lys Ser Gly Ala Ala Lys Ile Asn Glu Met Phe Ser Ile Ala Arg Arg Asn Ala Pro Ala Phe Val Phe Val Asp Glu Ile Asp Ala Ile Ala Gly Arg His Ala Arg Lys Asp Pro Arg Arg Arg Ala Thr Phe Glu Ala Leu Ile Ala Gln Leu Asp Gly Glu Lys Glu Lys

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Gln Ser Ala Arg Leu Gly Leu Thr Gln Leu Val Lys Lys Ile Gly Met 865 · 870 875 880

Val Asp Leu Pro Asp Asn Pro Asp Gly Glu Leu Ile Lys Tyr Arg Trp 885 890 895

Asp His Pro His Val Met Pro Ala Glu Met Ser Val Glu Val Ser Glu 900 905 910

Leu Phe Thr Arg Glu Leu Thr Arg Tyr Ile Glu Glu Thr Glu Glu Leu 915 920 925

Ala Met Asn Ala Leu Arg Ala Asn Arg His Ile Leu Asp Leu Ile Thr 930 935 940

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Lys Met Lys Asp Leu Ser Pro Leu Met Phe Glu Asp Phe Val Lys Pro 965 970 975

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								ttg Leu								1632
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gcc aaa tct atg gaa tgt cga gtt aag aag aaa gga aaa gtc ttc tgg Ala Lys Ser Met Glu Cys Arg Val Lys Lys Lys Gly Lys Val Phe Trp 770 775 780	2352
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Ser Gln Pro Ile Tyr Asn Ile Tyr Leu Asp Ser Leu Thr Lys Ile Gly Asn Leu Glu Lys Ala Gly Asp Val Phe Asn Glu Met Lys Asn Asn Gly 550 555 Thr Ile Asn Val Ser Ala Arg Ser Cys Asn Ser Leu Leu Lys Gly Tyr 570 Leu Asp Cys Gly Lys Gln Val Gln Ala Glu Arg Ile Tyr Asp Leu Met 580 585 Arg Met Lys Lys Tyr Glu Ile Glu Pro Pro Leu Met Glu Lys Leu Asp Tyr Ile Leu Ser Leu Lys Lys Glu Val Lys Lys Arg Pro Phe Ser 615 Met Lys Leu Ser Lys Asp Gln Arg Glu Val Leu Val Gly Leu Leu Leu 630 Gly Gly Leu Gln Ile Glu Ser Asp Lys Glu Lys Lys Ser His Met Ile 645 650 Lys Phe Glu Phe Arg Glu Asn Ser Gln Ala His Leu Val Leu Lys Gln 660 665 Asn Ile His Asp Gln Phe Arg Glu Trp Leu His Pro Leu Ser Asn Phe 675 680 Gln Glu Asp Ile Ile Pro Phe Glu Phe Tyr Ser Val Pro His Ser Tyr Phe Gly Phe Tyr Ala Glu His Tyr Trp Pro Lys Gly Gln Pro Glu Ile 710 715 Pro Lys Leu Ile His Arg Trp Leu Ser Pro His Ser Leu Ala Tyr Trp 725 730 735 Tyr Met Tyr Ser Gly Val Lys Thr Ser Ser Gly Asp Ile Ile Leu Arg 740 Leu Lys Gly Ser Leu Glu Gly Val Glu Lys Val Val Lys Ala Leu Gln 755 760 765

Ala Lys Ser Met Glu Cys Arg Val Lys Lys Lys Gly Lys Val Phe Trp 770 Ile Gly Leu Gln Gly Thr Asn Ser Ala Leu Phe Trp Lys Leu Ile Glu 790 795 Pro His Val Leu Glu Asn Leu Lys Glu His Leu Lys Pro Ala Ser Glu 805 810 Ser Leu Asp Asn Val Lys Glu Ala Glu Glu Gln Ser Ile Asn Phe Lys 820 825 830 Ser Asn Ser Asp His Ser Asp Asp Cys Val Asn Ser Glu Ala His Phe Tyr <210> 9 <211> 1449 <212> DNA <213> Arabidopsis thaliana <220> <221> CDS (1)..(1449) <222> <223> 8794 atg gtt gga tct ttg gaa tct gat caa act ctt tca atg gcc acc tta 48 Met Val Gly Ser Leu Glu Ser Asp Gln Thr Leu Ser Met Ala Thr Leu atc gaa aaa ctc gac atc tta tct gac gac ttc gat cca acc gcc gta 96 Ile Glu Lys Leu Asp Ile Leu Ser Asp Asp Phe Asp Pro Thr Ala Val gtc acc gaa ccg tta cct cct ccg gta act aat gga atc gga gct gat 144 Val Thr Glu Pro Leu Pro Pro Pro Val Thr Asn Gly Ile Gly Ala Asp 35 aaa gga gga gga gga gga gaa aga gag atg gtt ctc ggt agg aat ata 192 Lys Gly Gly Gly Gly Glu Arg Glu Met Val Leu Gly Arg Asn Ile 50

•

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	gat Asp														816
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		355 355														1104
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His Thr Thr Ser Leu Ala Val Thr Glu Pro Glu Val Asn Asp Glu Phe 65 70 75 80

Thr Gly Asp Lys Glu Ala Tyr Met Ala Ser Val Leu Ala Arg Tyr Arg 85 90 95

Lys Thr Leu Val Glu Arg Thr Lys Asn His Leu Gly Tyr Pro Tyr Asn 100 105 110

Leu Asp Phe Asp Tyr Gly Ala Leu Gly Gln Leu Gln His Phe Ser Ile 115 120 125

Asn Asn Leu Gly Asp Pro Phe Ile Glu Ser Asn Tyr Gly Val His Ser 130 135 140

Arg Pro Phe Glu Val Gly Val Leu Asp Trp Phe Ala Arg Leu Trp Glu 145 150 155 160

Ile Glu Arg Asp Asp Tyr Trp Gly Tyr Ile Thr Asn Cys Gly Thr Glu 165 170 175

Gly Asn Leu His Gly Ile Leu Val Gly Arg Glu Met Phe Pro Asp Gly 180 185 190

Ile Leu Tyr Ala Ser Arg Glu Ser His Tyr Ser Val Phe Lys Ala Ala 195 200 205

Arg Met Tyr Arg Met Glu Cys Glu Lys Val Asp Thr Leu Met Ser Gly 210 215 220

Glu Ile Asp Cys Asp Asp Leu Arg Lys Lys Leu Leu Ala Asn Lys Asp 225 230 235 240

Lys Pro Ala Ile Leu Asn Val Asn Ile Gly Thr Thr Val Lys Gly Ala 245 250 255

Val Asp Asp Leu Asp Leu Val Ile Lys Thr Leu Glu Glu Cys Gly Phe 260 265 270

Ser His Asp Arg Phe Tyr Ile His Cys Asp Gly Ala Leu Phe Gly Leu 275 280 285

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Pro Cys Gly Val Gln Ile Thr Arg Met Glu His Ile Lys Val Leu Ser 325 330 335

Ser Asn Val Glu Tyr Leu Ala Ser Arg Asp Ala Thr Ile Met Gly Ser 340 345 350

Arg Asn Gly His Ala Pro Leu Phe Leu Trp Tyr Thr Leu Asn Arg Lys 355 360 365

Gly Tyr Lys Gly Phe Gln Lys Glu Val Gln Lys Cys Leu Arg Asn Ala 370 380

His Tyr Leu Lys Asp Arg Leu Arg Glu Ala Gly Ile Ser Ala Met Leu 385 390 395 400

Asn Glu Leu Ser Ser Thr Val Val Phe Glu Arg Pro Lys Asp Glu Glu 405 410 415

Phe Val Arg Arg Trp Gln Leu Ala Cys Gln Gly Asp Ile Ala His Val 420 425 430

Val Val Met Pro Ser Val Thr Ile Glu Lys Leu Asp Asn Phe Leu Lys 435 440 445

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Ala Ile Asp Gly Ala Tyr Val Asp Lys Lys Cys Pro Phe Thr Gly Thr 50 55 60

Val Ser Ile Arg Gly Arg Ile Leu Ala Gly Thr Cys His Ser Ala Lys 65 70 75 80

Met Gln Arg Thr Ile Ile Val Arg Arg Asp Tyr Leu His Phe Val Lys 85 90 95

Lys Tyr Gln Arg Tyr Glu Lys Arg His Ser Asn Ile Pro Ala His Val 100 105 110

Ser Pro Cys Phe Arg Val Lys Glu Gly Asp His Ile Ile Gly Gln 115 120 125

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Tyr Lys Ala Gly Met Thr His Ile Val Arg Glu Val Glu Lys Pro Gly 50 55 60

Ser Lys Leu His Lys Lys Glu Thr Cys Glu Ala Val Thr Ile Ile Glu 65 70 75 80

Thr Pro Ala Met Val Val Val Gly Val Val Ala Tyr Val Lys Thr Pro 85 90 95

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Glu Val Arg Arg Arg Phe Tyr Lys Asn Trp Ala Lys Ser Lys Lys 115 120 125

Ala Phe Thr Gly Tyr Ala Lys Gln Tyr Asp Ser Glu Asp Gly Lys Lys 130 135 140

Gly Ile Gln Ala Gln Leu Glu Lys Met Lys Lys Tyr Ala Thr Val Ile 145 150 155 160

Arg Val Leu Ala His Thr Gln Ile Arg Lys Met Lys Gly Leu Lys Gln 165 170 175

Lys Lys Ala His Met Met Glu Ile Gln Ile Asn Gly Gly Thr Ile Ala 180 185 190

Gln Lys Val Asp Phe Ala Tyr Ser Phe Phe Glu Lys Gln Ile Pro Ile 195 200 205

Glu Ala Val Phe Gln Lys Asp Glu Met Ile Asp Ile Ile Gly Val Thr 210 215 220

Lys Gly Lys Gly Tyr Glu Gly Val Val Thr Arg Trp Gly Val Thr Arg 225 230 235 240

Leu Pro Arg Lys Thr His Arg Gly Leu Arg Lys Val Ala Cys Ile Gly 245 250 255

Ala Trp His Pro Ala Arg Val Ser Tyr Thr Val Ala Arg Ala Gly Gln 260 265 270

Asn Gly Tyr His His Arg Thr Glu Leu Asn Lys Lys Ile Tyr Arg Leu 275 280 285

Gly Lys Val Gly Thr Glu Ala His Thr Ala Met Thr Glu Tyr Asp Arg 290 295 300

Thr Glu Lys Asp Val Thr Pro Met Gly Gly Phe Pro His Tyr Gly Ile 305 310 315 320

Val Lys Asp Asp Tyr Leu Met Ile Lys Gly Cys Cys Val Gly Pro Lys 325 330 335

Lys Arg Val Val Thr Leu Arg Gln Ser Leu Leu Thr Gln Thr Ser Arg 340 345 350

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Asn Arg Val Thr Lys 385

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Tyr Val Lys Arg Thr Ser Lys Phe Met Ala His Asp Asp Lys Asp Ala 35 40 45

Cys Asn Ile Gly Asp Arg Val Lys Leu Asp Pro Ser Arg Pro Leu Ser 50 55 60

Lys Asn Lys His Trp Ile Val Ala Glu Ile Ile Lys Lys Ala Arg Ile 65 70 75 80

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gcc aat cag co Ala Asn Gln Le 290	ett act aac ata co eu Thr Asn Ile Le 295	tc aga gac gta ggc gaa eu Arg Asp Val Gly Glu 300	a gat gcg aga 912 1 Asp Ala Arg
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tca gat gaa ga Ser Asp Glu A	ac ata ttc gcc g sp Ile Phe Ala G 325	ga aaa gta act gat aaa ly Lys Val Thr Asp Lys 330	a tgg aga aac 1008 s Trp Arg Asn 335
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gct tca ttg c Ala Ser Leu L 370	eta ttg tac agg ag beu Leu Tyr Arg Ar 375	ga ata ctg gac gag at rg Ile Leu Asp Glu Ile 380	t gaa gcg aat 1152 e Glu Ala Asn
		ga gct tat gtg ggg aaa arg Ala Tyr Val Gly Lya 395	
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Pro Asp Pro Met Asn Asn Cys Gly Leu Val Arg Val Leu Glu Ser Ser

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Lys Gln Ile Pro Thr Trp Ser Ser Ser Phe Val Arg Asn Arg Ser Arg 50 55 60

Arg Ile Gly Val Val Ser Ser Ser Leu Val Ala Ser Pro Ser Gly Glu 65 70 75 80

Ile Ala Leu Ser Ser Glu Glu Lys Val Tyr Asn Val Val Leu Lys Gln 85 90 95

Ala Ala Leu Val Asn Lys Gln Leu Arg Ser Ser Ser Tyr Asp Leu Asp 100 105 110

Val Lys Lys Pro Gln Asp Val Val Leu Pro Gly Ser Leu Ser Leu Leu 115 120 125

Gly Glu Ala Tyr Asp Arg Cys Gly Glu Val Cys Ala Glu Tyr Pro Lys 130 135 140

Thr Phe Tyr Leu Gly Thr Leu Leu Met Thr Pro Glu Arg Arg Lys Ala 145 150 155 160

Ile Trp Ala Ile Tyr Val Trp Cys Arg Arg Thr Asp Glu Leu Val Asp 165 170 175

Gly Pro Asn Ala Ser His Ile Thr Pro Met Ala Leu Asp Arg Trp Glu 180 185 190

Ala Arg Leu Glu Asp Leu Phe Arg Gly Arg Pro Phe Asp Met Leu Asp 195 200 205

Ala Ala Leu Ala Asp Thr Val Ala Arg Tyr Pro Val Asp Ile Gln Pro 210 215 220

Phe Arg Asp Met Ile Glu Gly Met Arg Met Asp Leu Lys Lys Ser Arg 225 230 235 240

Tyr Gln Asn Phe Asp Asp Leu Tyr Leu Tyr Cys Tyr Tyr Val Ala Gly 245 250 255

Thr Val Gly Leu Met Ser Val Pro Val Met Gly Ile Asp Pro Lys Ser

260 265 270

Lys Ala Thr Thr Glu Ser Val Tyr Asn Ala Ala Leu Ala Leu Gly Ile 275 280 285

Ala Asn Gln Leu Thr Asn Ile Leu Arg Asp Val Gly Glu Asp Ala Arg 290 295 300

Arg Gly Arg Val Tyr Leu Pro Gln Asp Glu Leu Ala Gln Ala Gly Leu 305 310 315 320

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Phe Met Lys Met Gln Leu Lys Arg Ala Arg Met Phe Phe Asp Glu Ala 340 345 350

Glu Lys Gly Val Thr Glu Leu Ser Ala Ala Ser Arg Trp Pro Val Trp 355 360 365

Ala Ser Leu Leu Leu Tyr Arg Arg Ile Leu Asp Glu Ile Glu Ala Asn 370 380

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			act Thr													288
			gag Glu 100													336
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			tat Tyr 260									816
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Asp Leu Phe Lys Asn Trp Gly Glu Leu Asp Asp Ser Leu Phe Ser Val 50 55 60

Glu Arg Val Ser Gly Gly Ile Thr Asn Leu Leu Leu Lys Val Ser Val 65 70 75 80

Lys Glu Asp Thr Asn Lys Glu Val Ser Val Thr Val Arg Leu Tyr Gly
85 90 95

Pro Asn Thr Glu Tyr Val Ile Asn Arg Glu Arg Glu Ile Leu Ala Ile 100 105 110

Lys Tyr Leu Ser Ala Ala Gly Phe Gly Ala Lys Leu Leu Gly Gly Phe 115 120 125

Gly Asn Gly Met Val Gln Ser Phe Ile Asn Ala Arg Thr Leu Glu Pro 130 135 140

Ser Asp Met Arg Glu Pro Lys Ile Ala Ala Gln Ile Ala Arg Glu Leu 145 150 155 160

Gly Lys Phe His Lys Val Asp Ile Pro Gly Ser Lys Glu Pro Gln Leu 165 170 175

Trp Val Asp Ile Leu Lys Phe Tyr Glu Lys Ala Ser Thr Leu Thr Phe 180 185 190

Glu Glu Pro Asp Lys Gln Lys Leu Phe Glu Thr Ile Ser Phe Glu Glu 195 200 205

Leu His Lys Glu Ile Ile Glu Leu Arg Glu Phe Thr Gly Leu Leu Asn 210 215 220

Ala Pro Val Val Phe Ala His Asn Asp Leu Leu Ser Gly Asn Phe Met 225 230 235 240

Leu Asn Asp Glu Glu Glu Lys Leu Tyr Leu Ile Asp Phe Glu Tyr Gly 245 250 255

Ser Tyr Asn Tyr Arg Gly Phe Asp Ile Gly Asn His Phe Asn Glu Tyr 260 265 270

Ala Gly Tyr Asp Cys Asp Tyr Ser Leu Tyr Pro Ser Lys Glu Glu Gln 275 280 Tyr His Phe Ile Lys His Tyr Leu Gln Pro Asp Lys Pro Asp Glu Val 295 Ser Ile Ala Glu Val Glu Ser Val Phe Val Glu Thr Asp Ala Tyr Lys 305 310 315 320 Leu Ala Ser His Leu Tyr Trp Ala Ile Trp Ala Ile Ile Gln Ala Arg 325 330 Met Ser Pro Ile Glu Phe Glu Tyr Leu Gly Tyr Phe Phe Leu Arg Tyr 340 345 Asn Glu Tyr Lys Lys Gln Lys Pro Leu Thr Phe Ser Leu Val Thr Ser 360 355 His Leu Ser Ala Ser Leu 370 <210> 21 <211> 2283 <212> DNA <213> Arabidopsis thaliana <220> <221> CDS <222> (1)..(2283) <223> 13823 <400> 21 atg gcg atg act ccg gtc gcg tca tca tct cca gtt tca acc tgc aga 48 Met Ala Met Thr Pro Val Ala Ser Ser Pro Val Ser Thr Cys Arg 5 ctc ttt cgc tgc aat ctc ctc cct gat ctc tta cct aag cct ctg ttt 96 Leu Phe Arg Cys Asn Leu Leu Pro Asp Leu Leu Pro Lys Pro Leu Phe 20 ctc tcc ctc ccc aaa cga aac aga att gcc tcg tgc cgc ttc act gta 144

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Ser Thr Ser Asp Thr Asp Thr Glu Thr Asp Thr Thr Ser Tyr Gly Arg 70 75 80

Gln Phe Phe Pro Leu Ala Ala Val Val Gly Gln Glu Gly Ile Lys Thr 85 90 95

Ala Leu Leu Gly Ala Val Asp Arg Glu Ile Gly Gly Ile Ala Ile 100 105 110

Ser Gly Arg Arg Gly Thr Ala Lys Thr Val Met Ala Arg Gly Leu His 115 120 125

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Asp Pro Ala Cys Pro Asp Glu Trp Glu Asp Asp Leu Asp Glu Arg Ile 145 150 155 160

Glu Tyr Asn Ala Asp Asn Thr Ile Lys Thr Glu Ile Val Lys Ser Pro 165 170 175

Phe Ile Gln Ile Pro Leu Gly Val Thr Glu Asp Arg Leu Ile Gly Ser 180 185 190

- Val Asp Val Glu Glu Ser Val Lys Arg Gly Thr Thr Val Phe Gln Pro 195 200 205
- Gly Leu Leu Ala Glu Ala His Arg Gly Val Leu Tyr Val Asp Glu Ile 210 215 220
- Asn Leu Leu Asp Glu Gly Ile Ser Asn Leu Leu Leu Asn Val Leu Thr 225 230 235 235
- Asp Gly Val Asn Ile Val Glu Arg Glu Gly Ile Ser Phe Arg His Pro 245 250 255
- Cys Lys Pro Leu Leu Ile Ala Thr Tyr Asn Pro Glu Glu Gly Ala Val 260 265 270
- Arg Glu His Leu Leu Asp Arg Val Ala Ile Asn Leu Ser Ala Asp Leu 275 280 285
- Pro Met Ser Phe Glu Asp Arg Val Ala Ala Val Gly Ile Ala Thr Gln 290 295 300
- Phe Gln Glu Arg Cys Asn Glu Val Phe Arg Met Val Asn Glu Glu Thr 305 310 315
- Glu Thr Ala Lys Thr Gln Ile Ile Leu Ala Arg Glu Tyr Leu Lys Asp 325 330 335
- Val Lys Ile Ser Arg Glu Gln Leu Lys Tyr Leu Val Leu Glu Ala Val 340 345 350
- Arg Gly Gly Val Gln Gly His Arg Ala Glu Leu Tyr Ala Ala Arg Val 355 360 365
- Ala Lys Cys Leu Ala Ala Ile Glu Gly Arg Glu Lys Val Thr Ile Asp 370 375 380
- Asp Leu Arg Lys Ala Val Glu Leu Val Ile Leu Pro Arg Ser Ser Leu 385 390 395 400
- Asp Glu Thr Pro Pro Glu Gln Gln Asn Gln Pro Pro Pro Pro Pro Pro 405 410 415

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Ser Thr Asp Pro Glu Ser Ile Ala Pro Asp Ala Pro Arg Pro Thr Ser 675 680 685

Lys Glu Leu Lys Asp Glu Ile Leu Glu Val Ala Gly Lys Ile Tyr Lys 690 695 700

Ala Gly Met Ser Leu Leu Val Ile Asp Thr Glu Asn Lys Phe Val Ser 705 710 715 720

Thr Gly Phe Ala Lys Glu Ile Ala Arg Val Ala Gln Gly Lys Tyr Tyr 725 730 735

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gt	agg	cct	ato	aaa	aag	cct	gat	att	tat	aag	gct	ttt	999	gta	gac	1680

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ato Ile	aag Lys 770	Thr	agg Arg	cat His	ttc Phe	gag Glu 775	Gln	gcc Ala	ttg Leu	tcc Ser	tta Leu 780	gtc Val	tca Ser	cca Pro	tct Ser	2352
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Phe Ala Arg Leu Thr Arg Gln Val Leu Leu Leu Asn Val Arg Gln Val 50 55 60

Leu Asn Val Arg Asn Asn Lys Arg Val Lys Asp Glu Asp Glu Asp Asp 65 70 75 80

Asn Ile Gly Asp Glu Glu Gly Ser Ala Ser Gln Arg Lys Lys Gln Arg 85 90 95

Arg Val Asp Glu Lys Glu Glu Lys Leu Gln Arg Ala Glu Gln Ser His 100 105 110

Leu Arg Lys Arg Asn Met Glu Arg Ser Val Ser Ser Ser Pro Ser Ser 115 120 125

Ser Ser Ser Glu Asp Ser Gly Asp Val Ser Thr Ser Glu Asp Ala 130 135 140

Val Tyr Gly Glu Lys Leu Ser Pro Pro Arg Phe Asp Leu Ile Asn Asp Ser Leu Arg Asp Asn Tyr Ala Lys Leu Asn Ser Ser Ser Lys Lys Pro Ile Gly Ser Pro Ala Glu Lys Asn Val Glu Val Glu Thr Val Ser Asn Lys Gly Arg Ser Lys Leu Ala Thr Met Gly Ala Arg Lys Glu Ala Lys Val Ser Leu Ser Leu Ser Gly Ala Thr Gly Asn Gly Asp Leu Glu Val Glu Gly Thr Lys Gly Pro Thr Phe Lys Asp Phe Gly Gly Ile Lys Lys Ile Leu Asp Glu Leu Glu Met Asn Val Leu Phe Pro Ile Leu Asn Pro Glu Pro Phe Lys Lys Ile Gly Val Lys Pro Pro Ser Gly Ile Leu Phe His Gly Pro Pro Gly Cys Gly Lys Thr Lys Leu Ala Asn Ala Ile Ala Asn Glu Ala Gly Val Pro Phe Tyr Lys Ile Ser Ala Thr Glu Val Ile Ser Gly Val Ser Gly Ala Ser Glu Glu Asn Ile Arg Glu Leu Phe Ser Lys Ala Tyr Arg Thr Ala Pro Ser Ile Val Phe Ile Asp Glu Ile Asp Ala Ile Gly Ser Lys Arg Glu Asn Gln Gln Arg Glu Met Glu Lys Arg Ile Val Thr Gln Leu Leu Thr Cys Met Asp Gly Pro Gly Asn Lys Gly Asp Lys Asn Ala Pro Asp Ser Ser Ala Gly Phe Val Leu Val Ile Gly

Ala Thr Asn Arg Pro Asp Ala Leu Asp Pro Ala Leu Arg Arg Ser Gly Arg Phe Glu Thr Glu Ile Ala Leu Thr Ala Pro Asp Glu Asp Ala Arg Ala Glu Ile Leu Ser Val Val Ala Gln Lys Leu Arg Leu Glu Gly Pro Phe Asp Lys Lys Arg Ile Ala Arg Leu Thr Pro Gly Phe Val Gly Ala Asp Leu Glu Ser Val Ala Tyr Leu Ala Gly Arg Lys Ala Ile Lys Arg Ile Leu Asp Ser Arg Lys Ser Glu Gln Ser Gly Asp Gly Glu Asp Asp Lys Ser Trp Leu Arg Met Pro Trp Pro Glu Glu Glu Leu Glu Lys Leu Phe Val Lys Met Ser Asp Phe Glu Glu Ala Val Asn Leu Val Gln Ala Ser Leu Thr Arg Glu Gly Phe Ser Ile Val Pro Asp Val Lys Trp Asp Asp Val Gly Gly Leu Asp His Leu Arg Leu Gln Phe Asn Arg Tyr Ile Val Arg Pro Ile Lys Lys Pro Asp Ile Tyr Lys Ala Phe Gly Val Asp Leu Glu Thr Gly Phe Leu Leu Tyr Gly Pro Pro Gly Cys Gly Lys Thr Leu Ile Ala Lys Ala Ala Ala Asn Glu Ala Gly Ala Asn Phe Met His Ile Lys Gly Ala Glu Leu Leu Asn Lys Tyr Val Gly Glu Ser Glu Leu Ala Ile Arg Thr Leu Phe Gln Arg Ala Arg Thr Cys Ala Pro Cys Val

Ile Phe Phe Asp Glu Val Asp Ala Leu Thr Thr Ser Arg Gly Lys Glu 625 630 635 640

Gly Ala Trp Val Val Glu Arg Leu Leu Asn Gln Phe Leu Val Glu Leu 645 650 655

Asp Gly Glu Arg Arg Asn Val Tyr Val Ile Gly Ala Thr Asn Arg 660 665 670

Pro Asp Val Val Asp Pro Ala Phe Leu Arg Pro Gly Arg Phe Gly Asn 675 680 685

Leu Leu Tyr Val Pro Leu Pro Asn Ala Asp Glu Arg Ala Ser Ile Leu 690 695 700

Lys Ala Ile Ala Arg Lys Lys Pro Ile Asp Pro Ser Val Asp Leu Asp 705 710 715 720

Gly Ile Ala Lys Asn Asn Cys Glu Gly Phe Ser Gly Ala Asp Leu Ala 725 730 735

His Leu Val Gln Lys Ala Thr Phe Gln Ala Val Glu Glu Met Ile Gly 740 745 750

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Ile Lys Thr Arg His Phe Glu Gln Ala Leu Ser Leu Val Ser Pro Ser 770 775 780

Val Asn Lys Gln Gln Arg Arg His Tyr Asp Ala Leu Ser Thr Lys Leu 785 790 795 800

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	gca Ala															768
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- Gly Leu Asp Gln Asp Gly Met Ile Gly Lys Val Leu Val Lys Asn Pro 165 170 175
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- Val Met Asn Phe Pro Gln Leu Leu Cys Arg Asp Val Asn Lys Ile Leu 210 215 220
- Lys Pro Asn Tyr Asp Tyr Leu Lys Glu Cys Gly Phe Gly Asp Ser Gln 225 230 235
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- Lys Asn Ser Leu Gln Pro Arg Ile Arg Phe Leu Val Gln Val Met Gly 260 265 270
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336

aac aat gag cca gag tac gag gag tct tat agc ata gtc ata ctt ccg

Asn	Asn	Glu	Pro 100	Glu	Tyr	Glu	Glu	Ser 105	Tyr	Ser	Ile	Val	Ile 110	Leu	Pro.	
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	att Ile 130															432
	caa Gln															480
	ttg Leu															528
	tgg Trp															576
	acc Thr															624
	gga Gly 210															672
	gct Ala															720
	tat Tyr															768
	cat His															816
	atg Met															864
	aac Asn 290															912
	gga Gly					_					_	_			_	960
	acg Thr															1008
ttt	tca	cat	cag	agc	tta	aag	atg	gct	ttt	gag	atg	gct	cct	gct	gat	1056

Dho	Cor	uic	Gln	Cor	T.OU	Tarc	Mot-	מות	Dha	Glu	Met	Δla	Dro	Δla	Δsn	
PHE	Ser	птр	340	Ser	Беα	пуъ	Met	345	FIIC	Giu	Mec	AIG	350	діа	мор	
			gac Asp													1104
			aag Lys													1152
			aga Arg													1200
			agc Ser													1248
			ttc Phe 420													1296
			gat Asp													1344
			ctt Leu													1392
gac Asp 465			ctt Leu													1440
			gaa Glu													1488
			atg Met 500													1536
			tgt Cys													1584
			tct Ser													1632
			tct Ser													1680
			gaa Glu													1728
gtt	ccg	gat	gtt	att	gat	atc	agc	cac	atg	cgt	agc	aaa	gga	ctc	caa	1776

Val	Pro	Asp	Val 580	Ile	Asp	Ile	Ser	His 585	Met	Arg	Ser	Lys	Gly 590	Leu	Gln	
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		_			_					_	_	cag Gln		_		1872
												ata Ile				1920
												tct Ser				1968
												tct Ser				2016
		_	_	_								gct Ala 685	_	_	_	2064
												gag Glu				2112
-		-						_		-		gac Asp	_	_	_	2160
		_				_		_	_		_	gga Gly			_	2208
												cac His				2256
			_					_			_	aaa Lys 765	_		_	2304
	_				_	_		-				act Thr	_			2352
			ggt Gly									gat Asp	tga			2394

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Gly Lys Asp Tyr Val Ser Trp Asn Tyr Glu Lys Thr Gly Asn Pro Val 50 55 60

Tyr Leu His Ile Lys Gln Thr Arg Lys Ser Ile Pro Glu Asp Arg Pro 65 70 75 80

Leu Lys Lys Pro Thr Leu Leu Ala Ile Gly Val Asp Gly Gly Phe Asp 85 90 95

Asn Asn Glu Pro Glu Tyr Glu Glu Ser Tyr Ser Ile Val Ile Leu Pro 100 105 110

Asp Phe Val Ser Leu Pro Phe Pro Ser Val Glu Leu Pro Glu Lys Val

Arg Ile Ala Val Asp Thr Val Val Asn Ala Val Gly Ala Glu Arg Lys
130 135 140

Glu Gln Val Ala Ala Trp Thr Ala Glu Lys Lys Leu Ile Ser Glu His 145 150 155 160

Ala Leu Thr Leu Gln Gln Ile Lys Ser Gly Ile Val Ile Pro Pro Ser 165 170 175

Gly Trp Lys Cys Ser Lys Cys Asp Lys Thr Glu Asn Leu Trp Leu Asn 180 185 190

Leu Thr Asp Gly Met Ile Leu Cys Gly Arg Lys Asn Trp Asp Gly Thr
195 200 205

Gly Gly Asn Asn His Ala Val Glu His Tyr Lys Glu Thr Ala Tyr Pro Leu Ala Val Lys Leu Gly Thr Ile Thr Ala Asp Leu Glu Ala Ala Asp Val Tyr Ser Tyr Pro Glu Asp Asp Ser Val Leu Asp Pro Leu Leu Ala Glu His Leu Ala His Phe Gly Ile Asp Phe Ser Ser Met Gln Lys Thr Glu Met Thr Thr Ala Glu Arg Glu Leu Asp Gln Asn Thr Asn Phe Asp Trp Asn Arg Ile Gln Glu Ser Gly Lys Glu Leu Val Pro Val Phe Gly Pro Gly Tyr Thr Gly Leu Val Asn Leu Gly Asn Ser Cys Tyr Leu Ala Ala Thr Met Gln Ile Val Phe Ser Thr His Ser Phe Ile Ser Arg Tyr Phe Ser His Gln Ser Leu Lys Met Ala Phe Glu Met Ala Pro Ala Asp Pro Thr Leu Asp Leu Asn Met Gln Leu Thr Lys Leu Gly His Gly Leu Leu Ser Gly Lys Tyr Ser Met Pro Ala Thr Gln Lys Asp Ala Thr Thr Gly Asp Pro Arg Gln Glu Gly Ile Pro Pro Arg Met Phe Lys Asn Val Ile Ala Ala Ser His Ala Glu Phe Ser Ser Met Arg Gln Gln Asp Ala Leu Asp Phe Phe Leu His Leu Val Gly Lys Val Glu Arg Ala Ser Asn Thr Thr Pro Asp Leu Asp Pro Ser Arg Ser Phe Lys Phe Gly Ile Glu

Glu Lys Ile Leu Cys Pro Ser Gly Lys Val Gly Tyr Asn Lys Arg Glu Asp Cys Ile Leu Ser Leu Asn Ile Pro Leu His Glu Ala Thr Asn Lys Asp Glu Leu Glu Ala Phe His Lys Gln Lys Ala Gly Lys Gly Leu Glu Glu Asn Asp Met Arg Ser Ser Asp Glu Ile Val Arg Pro Arg Val Pro Leu Glu Ala Cys Leu Ala Asn Phe Ala Ser Ser Glu Pro Ile Glu Asp Tyr Tyr Ser Ser Ala Leu Lys Gly Met Thr Thr Ala Ile Lys Thr Thr Gly Leu Thr Ser Phe Pro Asp Tyr Leu Val Leu His Met Arg Lys Phe Val Met Glu Glu Gly Trp Val Pro Lys Leu Asp Val Tyr Ile Asp Val Pro Asp Val Ile Asp Ile Ser His Met Arg Ser Lys Gly Leu Gln Pro Gly Glu Glu Leu Pro Asp Gly Val Pro Glu Glu Val Met Glu Ser Ala Gln Pro Val Ala Asn Glu Glu Ile Val Ala Gln Leu Val Ser Met Gly Phe Ser Gln Leu His Cys Gln Lys Ala Ala Ile Asn Thr Ser Asn Ala Gly Val Glu Glu Ala Met Asn Trp Leu Leu Ser His Met Asp Asp Pro Asp Ile Asp Ala Pro Ile Ser His Gln Thr Ser Asp Ile Asp Gln Ser Ser Val Asp Thr Leu Leu Ser Phe Gly Phe Ala Glu Asp Val

Ala Arg Lys Ala Leu Lys Ala Ser Gly Gly Asp Ile Glu Lys Ala Thr 690 695 Asp Trp Val Phe Asn Asn Pro Asn Ala Ser Val Ser Asp Met Asp Val 710 715 Ser Ser Ser Asn Ser Ala Gln Thr Pro Ala Gln Ser Gly Leu Pro Asp 725 730 Gly Gly Gly Lys Tyr Lys Leu Phe Gly Ile Val Ser His Met Gly Thr 745 740 Ser Val His Cys Gly His Tyr Val Ala His Ile Leu Lys Glu Gly Arg Trp Val Ile Phe Asn Asp Asp Lys Val Gly Ile Ser Thr Asp Pro Pro 775 Lys Asp Met Gly Tyr Val Tyr Phe Phe Gln Arg Leu Asp 790 795 <210> 31 <211> 1068 <212> DNA <213> Arabidopsis thaliana <220> <221> CDS <222> (1)..(1068) <223> 15377 <400> 31 atg ata cta cgt cgt ttc atc tgc tac aac gct tct tca act gtt tca 48 Met Ile Leu Arg Arg Phe Ile Cys Tyr Asn Ala Ser Ser Thr Val Ser tct ata gct cca tca ccg aag aag cct ctc atc ttc tta ggc tct 96 Ser Ile Ala Pro Ser Pro Lys Lys Pro Leu Ile Phe Leu Gly Ser 25 20 cct cag gtc tcc gtg agt gtg ctt gaa gct ctt ttc aat gca tct aat 144 Pro Gln Val Ser Val Ser Val Leu Glu Ala Leu Phe Asn Ala Ser Asn 35

gct Ala	cca Pro 50	aac Asn	tct Ser	tcc Ser	ttc Phe	gag Glu 55	gtt Val	gca Ala	ggt Gly	att Ile	gtt Val 60	aca Thr	cag Gln	cct Pro	cca Pro	:	192
			gat Asp													2	240
			gat Asp													2	288
			gat Asp 100													:	336
			att Ile													3	384
			ccg Pro													4	132
			cgt Arg													· 4	180
			aca Thr													5	528
			gtg Val 180													Ę	576
			gaa Glu													6	524
			gaa Glu													€	572
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			gct Ala													7	768
			cgt Arg 260													8	316
			gat Asp													8	364

	_	tcc Ser		_	_	_		_	_	_		_	-	_	_	912
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Met 1	Ile		_	5			_	-	10					15		
Met 1 Ser	Ile	Leu	Pro 20	5 Ser	Pro	Lys	Lys	Lys 25	10 Pro	Leu	Ile	Phe	Leu 30	15 Gly	Ser	
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Met 1 Ser Pro	Ile Ile Gln Pro 50	Leu Ala Val 35	Pro 20 Ser	Ser Val Ser	Pro Ser Phe	Lys Val Glu 55	Lys Leu 40 Val	Lys 25 Glu Ala	10 Pro Ala Gly	Leu Leu Ile	Ile Phe Val 60	Phe Asn 45 Thr	Leu 30 Ala Gln	Gly Ser Pro	Ser Asn Pro	
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Glu Leu Cys Ile Thr Ala Ala Tyr Gly Asn Ile Leu Pro Thr Lys Phe Leu Lys Ile Pro Val His Gly Thr Val Asn Ile His Pro Ser Leu Leu Pro Leu Tyr Arg Gly Ala Ala Pro Val Gln Arg Ala Leu Gln Asp Gly Val Pro Glu Thr Gly Val Ser Leu Ala Phe Thr Val Arg Lys Leu Asp Ala Gly Pro Val Ile Ala Ser Lys Arg Phe Gln Val Asp Asp Leu Ile Lys Ala Pro Glu Leu Leu Ser Phe Leu Phe Ser Glu Gly Ser Asn Leu Leu Ile Arg Glu Leu Pro Ser Ile Phe Asp Gly Ser Ala Lys Ser Lys Ala Ala Pro Gln Asp Asp Ser Lys Ala Thr Leu Ala Pro Lys Ile Ala Pro Asp Glu Ala Trp Leu Ser Phe Asp Gln Glu Ala Phe Val Leu His Asn Lys Val Arg Ala Phe Ala Gly Trp Pro Gly Thr Arg Ala Lys Val Val Val Leu Asp Glu Lys Ser Gly Gln Gln Asn Val Leu Glu Leu Lys Ile Met Ser Thr Arg Val Cys Lys Asp Leu Glu Ile Gln Asp Ser Glu Gln Asp Tyr Val Thr Phe Lys Lys Gly Ser Leu Ile Phe Pro Cys Arg Gly Gly Thr Ala Leu Glu Val Leu Glu Val Gln Leu Pro Gly Lys Lys Ala Ile Asn Ala Ala Phe Trp Asn Gly Leu Arg Gly Gln Lys Leu

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	cgg Arg															576
	cct Pro															624
	acc Thr 210															672
	cat His		_	_				_		_		-				720
	cgt Arg															768
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	att Ile															864
	aag Lys 290															912
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	aag Lys															1008
	gag Glu															1056
	act Thr															1104
gcc	cca	cct	cca	aaa	ttt	gtt	aaa	gtc	acc	atg	tct	aag	agg	gtt	cct	1152

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aaa Lys	agg Arg	gaa Glu	gaa Glu	gct Ala 405	ttg Leu	cga Arg	gct Ala	agc Ser	ctc Leu 410	gtt Val	aaa Lys	gag Glu	gag Glu	gaa Glu 415	aca Thr	1248
				gga Gly												1296
gat Asp	acc Thr	aag Lys 435	act Thr	act Thr	cac His	gat Asp	gtt Val 440	gtt Val	ggt Gly	tct Ser	cac His	ggg Gly 445	cct Pro	gca Ala	tat Tyr	1344
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gag Glu	att Ile	att Ile	aat Asn	cca Pro 485	gat Asp	gac Asp	tat Tyr	gtg Val	atc Ile 490	aag Lys	gat Asp	gaa Glu	gac Asp	atg Met 495	gac Asp	1488
cga Arg	gga Gly	gca Ala	atg Met 500	cat His	aac Asn	gga Gly	ggt Gly	gat Asp 505	gtg Val	gac Asp	gga Gly	agg Arg	ctt Leu 510	gat Asp	gag Glu	1536
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gaa Glu 545	ggt Gly	cgg Arg	tca Ser	gat Asp	ggc Gly 550	cgc Arg	tca Ser	atc Ile	aag Lys	tca Ser 555	atg Met	att Ile	gcg Ala	cat His	gtt Val 560	1680
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cat His	ttg Leu	aag Lys	caa Gln 580	cac His	tgc Cys	ttg Leu	aac Asn	aac Asn 585	atc Ile	tgt Cys	cca Pro	cac His	gtg Val 590	tat Tyr	gct Ala	1776
cct Pro	caa Gln	ata Ile 595	gag Glu	gaa Glu	acg Thr	gtc Val	gat Asp 600	gtg Val	act Thr	tct Ser	gat Asp	tta Leu 605	tgt Cys	gct Ala	tac Tyr	1824
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gag Glu	agg Arg	gac Asp	atg Met	agg Arg 645	tct Ser	cta Leu	cta Leu	ccg Pro	atg Met 650	cca Pro	ggt Gly	gct Ala	gct Ala	tcg Ser 655	cca Pro	1968
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Pro Arg Val Ala Ser Thr Ile Asp Ala Val Leu Ser His Pro Asp 50 55 60

Thr Leu His Ile Gly Ala Leu Pro Tyr Ala Met Lys Gln Leu Gly Leu 65 70 75 80

Ser Ala Pro Val Tyr Ala Thr Glu Pro Val His Arg Leu Gly Leu Leu 85 90 95

Thr Met Tyr Asp Gln Phe Leu Ser Arg Lys Gln Val Ser Asp Phe Asp 100 105 110

Leu Phe Thr Leu Asp Asp Ile Asp Ser Ala Phe Gln Asn Val Ile Arg 115 120 125

Leu Thr Tyr Ser Gln Asn Tyr His Leu Ser Gly Lys Gly Glu Gly Ile 130 135 140

Val Ile Ala Pro His Val Ala Gly His Met Leu Gly Gly Ser Ile Trp 145 150 155 160

Arg Ile Thr Lys Asp Gly Glu Asp Val Ile Tyr Ala Val Asp Tyr Asn 165 170 175

His Arg Lys Glu Arg His Leu Asn Gly Thr Val Leu Gln Ser Phe Val

Arg Pro Ala Val Leu Ile Thr Asp Ala Tyr His Ala Leu Tyr Thr Asn 195 200 205

Gln Thr Ala Arg Gln Gln Arg Asp Lys Glu Phe Leu Asp Thr Ile Ser 210 215 220

Lys His Leu Glu Val Gly Gly Asn Val Leu Leu Pro Val Asp Thr Ala 225 230 235 240

Gly Arg Val Leu Glu Leu Leu Leu Ile Leu Glu Gln His Trp Ser Gln 245 250 255

Arg Gly Phe Ser Phe Pro Ile Tyr Phe Leu Thr Tyr Val Ser Ser Ser 260 265 270

Thr Ile Asp Tyr Val Lys Ser Phe Leu Glu Trp Met Ser Asp Ser Ile 275 280 285

Ser Lys Ser Phe Glu Thr Ser Arg Asp Asn Ala Phe Leu Leu Arg His Val Thr Leu Leu Ile Asn Lys Thr Asp Leu Asp Asn Ala Pro Pro Gly Pro Lys Val Val Leu Ala Ser Met Ala Ser Leu Glu Ala Gly Phe Ala Arg Glu Ile Phe Val Glu Trp Ala Asn Asp Pro Arg Asn Leu Val Leu Phe Thr Glu Thr Gly Gln Phe Gly Thr Leu Ala Arg Met Leu Gln Ser Ala Pro Pro Pro Lys Phe Val Lys Val Thr Met Ser Lys Arg Val Pro Leu Ala Gly Glu Glu Leu Ile Ala Tyr Glu Glu Glu Gln Asn Arg Leu Lys Arg Glu Glu Ala Leu Arg Ala Ser Leu Val Lys Glu Glu Glu Thr Lys Ala Ser His Gly Ser Asp Asp Asn Ser Ser Glu Pro Met Ile Ile Asp Thr Lys Thr Thr His Asp Val Val Gly Ser His Gly Pro Ala Tyr Lys Asp Ile Leu Ile Asp Gly Phe Val Pro Pro Ser Ser Ser Val Ala Pro Met Phe Pro Tyr Tyr Asp Asn Thr Ser Glu Trp Asp Asp Phe Gly Glu Ile Ile Asn Pro Asp Asp Tyr Val Ile Lys Asp Glu Asp Met Asp Arg Gly Ala Met His Asn Gly Gly Asp Val Asp Gly Arg Leu Asp Glu Ala Thr Ala Ser Leu Met Leu Asp Thr Arg Pro Ser Lys Val Met Ser

Asn Glu Leu Ile Val Thr Val Ser Cys Ser Leu Val Lys Met Asp Tyr 530 540

Glu Gly Arg Ser Asp Gly Arg Ser Ile Lys Ser Met Ile Ala His Val 545 550 555 560

Ser Pro Leu Lys Leu Val Leu Val His Ala Ile Ala Glu Ala Thr Glu 565 570 575

His Leu Lys Gln His Cys Leu Asn Asn Ile Cys Pro His Val Tyr Ala 580 585 590

Pro Gln Ile Glu Glu Thr Val Asp Val Thr Ser Asp Leu Cys Ala Tyr 595 600 605

Lys Val Gln Leu Ser Glu Lys Leu Met Ser Asn Val Ile Phe Lys Lys 610 615 620

Leu Gly Asp Ser Glu Val Ala Trp Val Asp Ser Glu Val Gly Lys Thr 625 630 635 640

Glu Arg Asp Met Arg Ser Leu Leu Pro Met Pro Gly Ala Ala Ser Pro 645 650 655

His Lys Pro Val Leu Val Gly Asp Leu Lys Ile Ala Asp Phe Lys Gln 660 665 670

Phe Leu Ser Ser Lys Gly Val Gln Val Glu Phe Ala Gly Gly Gly Ala 675 680 685

Leu Arg Cys Gly Glu Tyr Val Thr Leu Arg Lys Val Gly Pro Thr Gly 690 695 700

Gln Lys Gly Gly Ala Ser Gly Pro Gln Gln Ile Leu Ile Glu Gly Pro 705 710 715 720

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cgg Arg	acg Thr 50	atc Ile	atc Ile	act Thr	ttc Phe	cag Gln 55	aaa Lys	att Ile	tca Ser	acc Thr	60 GJÀ 333	att Ile	gtt Val	cct Pro	cca Pro	192
cca Pro 65	tcg Ser	gct Ala	tca Ser	tca Ser	tct Ser 70	ccg Pro	tcg Ser	agc Ser	tat Tyr	gga Gly 75	gac Asp	ctt Leu	caa Gln	cca Pro	atc Ile 80	240
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gta Val	caa Gln	gag Glu	cca Pro 100	aag Lys	gct Ala	aaa Lys	tac Tyr	gag Glu 105	cag Gln	ctt Leu	atg Met	ttc Phe	tac Tyr 110	gly aaa	aag Lys	336
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gaa Glu	gga Gly 130	tgt Cys	gtt Val	tct Ser	cag Gln	gtt Val 135	tgg Trp	gtt Val	agg Arg	gct Ala	ttc Phe 140	ttt Phe	gat Asp	gag Glu	gaa Glu	432
cgt Arg 145	aat Asn	gtt Val	gtg Val	tat Tyr	gaa Glu 150	gct Ala	gat Asp	tct Ser	gat Asp	tcg Ser 155	gtt Val	ctc Leu	act Thr	aaa Lys	999 Gly 160	480
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atg cag Met Gli 210	Lys														672
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Val Gln Glu Pro Lys Ala Lys Tyr Glu Gln Leu Met Phe Tyr Gly Lys
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Glu Gly Cys Val Ser Gln Val Trp Val Arg Ala Phe Phe Asp Glu Glu 130 135 140

Arg Asn Val Val Tyr Glu Ala Asp Ser Asp Ser Val Leu Thr Lys Gly 145 150 155 160

Leu Ala Ala Leu Leu Val Lys Gly Leu Ser Gly Arg Pro Val Pro Glu 165 170 175

Ile Leu Arg Ile Thr Pro Asp Phe Ala Val Leu Leu Gly Leu Gln Gln 180 185 190

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Leu Val Glu Asp Leu Gly Thr Glu Lys Ile Asp Asp Ser Glu Ser Gly 260 265 270

Ser Asn Val Val Ala Leu Gly Ser Arg Gly Met Arg Ile Arg Glu Lys 275 280 285

Leu Glu Lys Glu Leu Asp Pro Val Glu Leu Glu Val Glu Asp Val Ser 290 295 300

Tyr Gln His Ala Gly His Ala Ala Val Arg Gly Ser Ala Gly Asp Asp 305 310 315 320

Gly Glu Thr His Phe Asn Leu Arg Ile Val Ser Asp Ala Phe Gln Gly 325 330 335

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act Thr	ttg Leu	gag Glu 115	gga Gly	cat His	gaa Glu	aac Asn	gaa Glu 120	gtc Val	aaa Lys	agt Ser	gta Val	tca Ser 125	tgg Trp	aat Asn	gca Ala	384
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Cys Lys Thr Val Leu Glu Glu Thr His Thr Arg Thr Val Arg Ser Cys Ala Trp Ser Pro Ser Gly Gln Leu Leu Ala Thr Ala Ser Phe Asp Gly Thr Thr Gly Ile Trp Lys Asn Tyr Gly Ser Glu Phe Glu Cys Ile Ser Thr Leu Glu Gly His Glu Asn Glu Val Lys Ser Val Ser Trp Asn Ala Ser Gly Ser Cys Leu Ala Thr Cys Ser Arg Asp Lys Ser Val Trp Ile Trp Glu Val Leu Glu Gly Asn Glu Tyr Asp Cys Ala Ala Val Leu Thr Gly His Thr Gln Asp Val Lys Met Val Gln Trp His Pro Thr Met Asp Val Leu Phe Ser Cys Ser Tyr Asp Asn Thr Ile Lys Val Trp Trp Ser Glu Asp Asp Gly Glu Tyr Gln Cys Val Gln Thr Leu Gly Glu Ser Asn Asn Gly His Ser Ser Thr Val Trp Ser Ile Ser Phe Asn Ala Ala Gly Asp Lys Met Val Thr Cys Ser Asp Asp Leu Thr Leu Lys Ile Trp Gly Thr Asp Ile Ala Lys Met Gln Ser Gly Glu Glu Tyr Ala Pro Trp Ile His Leu Cys Thr Leu Ser Gly Tyr His Asp Arg Thr Ile Tyr Ser Ala His Trp Ser Arg Asp Asp Ile Ile Ala Ser Gly Ala Gly Asp Asn Ala Ile Arg Leu Phe Val Asp Ser Lys His Asp Ser Val Asp Gly Pro

Ser Tyr Asn Leu Leu Leu Lys Lys Asn Lys Ala His Glu Asn Asp Val 315 305 310 Asn Ser Val Gln Trp Ser Pro Gly Glu Gly Asn Arg Leu Leu Ala Ser 325 Ala Ser Asp Asp Gly Met Val Lys Ile Trp Gln Leu Ala Thr Lys Pro 345 340 <210> 39 <211> 942 <212> DNA <213> Arabidopsis thaliana <220> <221> CDS <222> (1)..(924) <223> atg cta agc ttg aga tat tca tta cct tat ctt ctt caa aca agg 48 Met Leu Ser Leu Arg Tyr Ser Leu Pro Tyr Leu Leu Gln Thr Arg 10 gaa tca tca act aag ctc ttc acc aaa aag cct aac aat gtt gtg gtt 96 Glu Ser Ser Thr Lys Leu Phe Thr Lys Lys Pro Asn Asn Val Val Val 20 tgt gcg gcg aga ggt cca aga cct cgg tct cct cgt gta tgg aaa aca 144 Cys Ala Ala Arg Gly Pro Arg Pro Arg Ser Pro Arg Val Trp Lys Thr 35 agg aag agg att gga act atc tct aaa gct gcc aaa atg att gct tgt Arg Lys Arg Ile Gly Thr Ile Ser Lys Ala Ala Lys Met Ile Ala Cys 50 ata aaa gga ttg tcg aat gtt aaa gaa gat tat gga gcg ctt gat 240 Ile Lys Gly Leu Ser Asn Val Lys Glu Glu Val Tyr Gly Ala Leu Asp tcc ttc att gct tgg gaa tta gag ttc cct ctt gtt ata gtt aag aag 288 Ser Phe Ile Ala Trp Glu Leu Glu Phe Pro Leu Val Ile Val Lys Lys Ala Leu Val Ile Leu Glu Asp Glu Lys Glu Trp Lys Lys Ile Ile Gln

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Ser Phe Ile Ala Trp Glu Leu Glu Phe Pro Leu Val Ile Val Lys Lys 85 90 95

Ala Leu Val Ile Leu Glu Asp Glu Lys Glu Trp Lys Lys Ile Ile Gln 100 105 110

Val Thr Lys Trp Met Leu Ser Lys Gly Gln Gly Arg Thr Met Gly Thr 115 120 125

Tyr Phe Ser Leu Leu Asn Ala Leu Ala Glu Asp Asn Arg Leu Asp Glu 130 135 140

Ala Glu Glu Leu Trp Asn Lys Leu Phe Met Glu His Leu Glu Gly Thr 145 150 155 160

Pro Arg Lys Phe Phe Asn Lys Met Ile Ser Ile Tyr Tyr Lys Arg Asp 165 170 175

Met His Gln Lys Leu Phe Glu Val Phe Ala Asp Met Glu Glu Leu Gly 180 185 190

Val Lys Pro Asn Val Ala Ile Val Ser Met Val Gly Lys Val Phe Val
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275 280 285

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Asp Ala Asp Leu Gln Leu Met Arg Asp Arg Ala Asn Ser Val Lys Asn 50 55 60

Leu Ala Ser Thr Phe Asp Arg Glu Ile Glu Asn Phe Leu Asn Asn Ser 65 70 75 80

Ala Arg Ser Ala Phe Pro Val Gly Ser Pro Ser Ala Ser Ser Phe Ser 85 90 95

Asn Glu Ile Gly Ile Met Lys Lys Leu Gln Pro Lys Ile Ser Glu Phe 100 105 110

Arg Arg Val Tyr Ser Ala Pro Glu Ile Ser Arg Lys Val Met Glu Arg 115 120 125

Trp Gly Pro Ala Arg Ala Lys Leu Gly Met Asp Leu Ser Ala Ile Lys

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Lys Ala Ile Val Ser Glu Met Glu Leu Asp Glu Arg Gln Gly Val Leu 145 150 155 160

Glu Met Ser Arg Leu Arg Arg Arg Arg Asn Ser Asp Arg Val Arg Phe 165 170 175

Thr Glu Phe Phe Ala Glu Ala Glu Arg Asp Gly Glu Ala Tyr Phe Gly 180 185 190

Asp Trp Glu Pro Ile Arg Ser Leu Lys Ser Arg Phe Lys Glu Phe Glu
195 200 205

Lys Arg Ser Ser Leu Glu Ile Leu Ser Gly Phe Lys Asn Ser Glu Phe 210 215 220

Val Glu Lys Leu Lys Thr Ser Phe Lys Ser Ile Tyr Lys Glu Thr Asp 225 230 235 240

Glu Ala Lys Asp Val Pro Pro Leu Asp Val Pro Glu Leu Leu Ala Cys 245 250 255

Leu Val Arg Gln Ser Glu Pro Phe Leu Asp Gln Ile Gly Val Arg Lys 260 265 270

Asp Thr Cys Asp Arg Ile Val Glu Ser Leu Cys Lys Cys Lys Ser Gln 275 280 285

Gln Leu Trp Arg Leu Pro Ser Ala Gln Ala Ser Asp Leu Ile Glu Asn 290 295 300

Asp Asn His Gly Val Asp Leu Asp Met Arg Ile Ala Ser Val Leu Gln 305 310 315

Ser Thr Gly His His Tyr Asp Gly Gly Phe Trp Thr Asp Phe Val Lys 325 330 335

Pro Glu Thr Pro Glu Asn Lys Arg His Val Ala Ile Val Thr Thr Ala 340 345 350

Ser Leu Pro Trp Met Thr Gly Thr Ala Val Asn Pro Leu Phe Arg Ala 355 360 365

Ala Tyr Leu Ala Lys Ala Ala Lys Gln Ser Val Thr Leu Val Val Pro

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Trp Leu Cys Glu Ser Asp Gln Glu Leu Val Tyr Pro Asn Asn Leu Thr 385 390 395 400

Phe Ser Ser Pro Glu Glu Gln Glu Ser Tyr Ile Arg Lys Trp Leu Glu 405 410 415

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Lys Phe Ser Lys Glu Arg Arg Ser Ile Phe Pro Ala Gly Asp Thr Ser 435 440 445

Gln Phe Ile Ser Ser Lys Asp Ala Asp Ile Ala Ile Leu Glu Glu Pro 450 455 460

Glu His Leu Asn Trp Tyr Tyr His Gly Lys Arg Trp Thr Asp Lys Phe 465 470 475 480

Asn His Val Val Gly Ile Val His Thr Asn Tyr Leu Glu Tyr Ile Lys 485 490 495

Arg Glu Lys Asn Gly Ala Leu Gln Ala Phe Phe Val Asn His Val Asn 500 505 510

Asn Trp Val Thr Arg Ala Tyr Cys Asp Lys Val Leu Arg Leu Ser Ala 515 520 525

Ala Thr Gln Asp Leu Pro Lys Ser Val Val Cys Asn Val His Gly Val 530 535 540

Asn Pro Lys Phe Leu Met Ile Gly Glu Lys Ile Ala Glu Glu Arg Ser 545 550 555

Arg Gly Glu Gln Ala Phe Ser Lys Gly Ala Tyr Phe Leu Gly Lys Met 565 570 575

Val Trp Ala Lys Gly Tyr Arg Glu Leu Ile Asp Leu Met Ala Lys His 580 585 590

Lys Ser Glu Leu Gly Ser Phe Asn Leu Asp Val Tyr Gly Asn Gly Glu
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Asp Ala Val Glu Val Gln Arg Ala Ala Lys Lys His Asp Leu Asn Leu

610 615 620

Asn Phe Leu Lys Gly Arg Asp His Ala Asp Asp Ala Leu His Lys Tyr 625 630 635 640

Lys Val Phe Ile Asn Pro Ser Ile Ser Asp Val Leu Cys Thr Ala Thr 645 650 655

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Ser Asn Glu Phe Phe Arg Ser Phe Pro Asn Cys Leu Thr Tyr Lys Thr 675 680 685

Ser Glu Asp Phe Val Ser Lys Val Gln Glu Ala Met Thr Lys Glu Pro 690 695 700

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Thr Gln Arg Phe Met Glu Tyr Ser Asp Leu Asp Lys Ile Leu Asn Asn 725 730 735

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Asn Glu Val Val Asp Gly Gly Leu Ala Phe Ser His Tyr Val Leu Thr 755 760 765

Gly Asn Asp Phe Leu Arg Leu Cys Thr Gly Ala Thr Pro Arg Thr Lys 770 775 780

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Ser Leu Asp Asn Asp Gly Asp Ser Ser Ser Ala Asp Cys Met His Glu 50 55 60

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Val Glu Thr Arg Thr Arg Phe Pro Glu Leu Val Ile His Glu Glu Lys 225 230 235

Arg Val Arg Phe Val Val Val Asn Gly Leu Asp Ile Val Glu Lys Pro 245 250 255

Ser Asp Leu Pro Ile Glu Glu Ala Glu Trp Phe Lys Arg Leu Thr Gly 260 265 270

Arg Asn Glu Val Ala Ile Ser Ala Arg Asp Tyr Lys Phe Tyr Cys Pro 275 280 285

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Lys His His Met Ser Ser Leu Ser His Gln Phe His Gln Ser Ile His 340 345 350

Gln Ser His Gln His His Gln Ser Ile Tyr Gln Ser Gln His Ala Ala 355 360 365

Thr His Tyr Pro Ser Gln Asn His Gln Cys Asp Pro Glu Leu Ser His 370 375 380

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130

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- Ile Glu Asp Gln Asn Ser Ile Ile Ile Ile Ile His Ala Thr Asn Asn 65 70 75 80
- Cys Leu Gln Arg Cys Pro Ser Val Thr Lys Glu Gln Trp Ala Val Pro 85 90 95
- Ala Ile Leu Ser Ser Leu Lys Met Glu Glu Asn Leu Leu Ala Gln Glu 100 105 110
- Arg Ala Cys Val Phe Leu Ser Leu Leu Leu His Asn Phe Ser Met Val
- His Thr Thr Lys Thr Gly Asn Thr Leu Asn Val Asp Ser Phe Ser Cys 130 135 140
- Leu Asp Ser Phe Ser Lys His Ile Arg Gly Gly Met Ala Asp Thr Glu 145 150 155 160
- Ala Gly Val Met Leu Ser Gly Phe Ser Glu Glu Leu Leu Cys Leu Leu 165 170 175
- Gln Asp Leu Leu Ser Gly Gln Arg Val Leu Phe Ser Val Lys Ser Ser 180 185 190
- Glu Thr Cys Glu Ser Asp Leu Ser Ile Pro Val Thr Leu Asn Gly Glu 195 200 205
- Asn Val Ala Leu Val Asn Lys Ile Ala Leu Thr Asp Gln Leu Val Ala 210 215 220
- Gly Ser Ala Ile Leu Ala Ala Ile Cys Thr Ala Leu Asp Arg Ile Gly 225 230 235 240
- Tyr Ile Cys Glu Ala Ser Phe Glu Ile Leu His Lys Tyr Ser His Glu 245 250 255
- Lys Thr Ser Val Leu Leu Thr Ile Leu His Val Phe Ala Tyr Ile Ala 260 265 270

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Ser Asn Leu Arg Val Lys Leu Ser Ala Phe Leu Gln Cys Glu Thr Thr 450 455 460

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Lys Thr Leu Gln Leu Lys Phe Pro Ile Asp Phe Gln Asp Lys Thr Thr 485 490 495

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H1S 225	Asp	GIU	Arg	Pne	230	TTE	PIO	Arg	vai	235	AIU	Val	Val	014	240	
tgc	att	cag	ggc	ggc Gly	gtt	gaa	ata	CCC	gaa	ttt	cca	gca	aaa	agg	aga Arg	768
Cys	TTE	GIN	GIĀ	245	vai	GIU	116	PLO	250	FIIC	FIO	AIG	מעם	255		
aga	aaa	ccg	att	atc	cgc	att	ggc	aaa	agc	gag	ttt	gtt	gat	gca	gat	816
Arg	гув	Pro	260	Ile	Arg	тте	GTĀ	ду5 265	per	GIU	FIIC	Vai	270	ALU	nop	
gaa	act	gaa	ttg	cct Pro	gat	cca	gag	cct	cag	cct	cct	cca	gtg	cca	ttg	864
GIU	Thr	275	ьец	PIO	Авр	PIO	280	FIO	GIII	FLO	110	285	741			
tta	act	gag	tta	cct Pro	gtc	tca	gag	atc	act	ccc	cca	tct	agc	gaa	gaa Glu	912
Leu	7nr 290	GIU	Leu	Pro	vai	295	GIU	TIE	1111	PLO	300	Ser	BCI.	O.L.	OIU	
gaa	aca	gtc	tcc	tta Leu	gcc	gaa	gaa	aca	tta	cag	gcc	tgg	gaa	gaa	atg	960
305	Thr	vai	ser	ьeu	310	GIU	GIU	TIIT	пеп	315	AIA	TTP	GIU	OLU	320	
aga	gca	gga	gcc	aaa	aag	ctg	atg	agg	atg	tac	agg	gtt	agg	gtc	tgt	1008
Arg	А1а	GIA	AIa	Lys 325	тув	ьeu	мес	Arg	330	TÄT	AIG	Vai	AL 9	335	Cyb	
a aa	tac	tgt	cca	gag	gtt	cac	gta	ggt	cca	acg	gga	cac His	aag	gcc Ala	cag Gln	1056
GТĀ	туr	сув	9ro 340		val	uis	val	345	LTO.	TIT	ary	11113	350			
aac	tgt	ggt	gca	ttc	aag	cac	caa	cag	cgg	aat	ggc	cag	cat	ggt	tgg	1104

Asn Cys G	ly Ala	Phe Ly	s His	Gln 360	Gln	Arg	Asn	Gly	Gln 365	His	Gly	Trp	
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gtt cct g Val Pro A 385	gat gtg Asp Val	aat gg Asn G	ly Pro	ccg Pro	atg Met	cag Gln	cga Arg 395	gag Glu _.	cta Leu	cga Arg	agc Ser	ttc Phe 400	1200
tac ggg o	caa gca 31n Ala	cct go Pro A 405	ct gtt la Val	gtg Val	gag Glu	ata Ile 410	tgt Cys	gct Ala	cag Gln	gct Ala	ggc Gly 415	gct Ala	1248
gtt gta d Val Val E	ect gag Pro Glu 420	cat to	at aga yr Arg	gct Ala	aca Thr 425	atg Met	aga Arg	ctg Leu	gag Glu	gtt Val 430	gga Gly	att Ile	1296
cct tcg a	agt gtg Ser Val 435	aaa g Lys G	aa gct lu Ala	gag Glu 440	atg Met	gtt Val	gtt Val	tga					1332
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	rT												
	rabidop	sis th	aliana										
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1	Ile Thr	Tyr S 5	er Ala	Ile	Ser	Ser 10	Ser	Thr	Val	Ser	Gly 15	Phe	
		5				10					1.5		
Ser Pro	Lys Ser 20	5 Val P	ro Phe	Ala	Ile 25	10 His	Ser	Val	Thr	Arg 30	Arg	Gln	
Ser Pro	Lys Ser 20 Asn Pro 35	Val P	ro Phe	Ala Tyr 40	Ile 25 Arg	His	Ser	Val Phe	Thr Ser 45	Arg 30 Pro	Arg Ser	Gln Leu	

Pro Gln Asn Glu Asp Leu Pro Lys Gln Tyr Thr Arg Arg Glu Lys Lys 85 90 95

Pro Phe Pro Val Pro Ile Val Asp Leu Arg Arg Ala Ala Arg Glu Arg Val Lys Asn Asn Lys Asp Lys Pro Lys Arg Pro Leu Pro Pro Pro Lys Asn Gly Met Val Val Lys Ser Leu Val Pro Leu Ala Tyr Lys Val Tyr Asn Ala Arg Ile Arg Leu Ile Asn Asn Leu His Arg Leu Met Lys Val Val Arg Val Asn Ala Cys Gly Trp Cys Asn Glu Ile His Val Gly Pro Tyr Gly His Pro Phe Lys Ser Cys Lys Gly Pro Asn Thr Ser Gln Arg Lys Gly Leu His Glu Trp Thr Asn Ser Val Ile Glu Asp Val Ile Val Pro Leu Glu Ala Tyr His Leu Phe Asp Arg Leu Gly Lys Arg Ile Arg His Asp Glu Arg Phe Ser Ile Pro Arg Val Pro Ala Val Val Glu Leu Cys Ile Gln Gly Gly Val Glu Ile Pro Glu Phe Pro Ala Lys Arg Arg Arg Lys Pro Ile Ile Arg Ile Gly Lys Ser Glu Phe Val Asp Ala Asp Glu Thr Glu Leu Pro Asp Pro Glu Pro Gln Pro Pro Pro Val Pro Leu Leu Thr Glu Leu Pro Val Ser Glu Ile Thr Pro Pro Ser Ser Glu Glu Glu Thr Val Ser Leu Ala Glu Glu Thr Leu Gln Ala Trp Glu Glu Met Arg Ala Gly Ala Lys Lys Leu Met Arg Met Tyr Arg Val Arg Val Cys

Gly Tyr Cys Pro Glu Val His Val Gly Pro Thr Gly His Lys Ala Gln 340 345 350

Asn Cys Gly Ala Phe Lys His Gln Gln Arg Asn Gly Gln His Gly Trp 355 360 365

Gln Ser Ala Val Leu Asp Asp Leu Ile Pro Pro Arg Tyr Val Trp His 370 375 380

Val Pro Asp Val Asn Gly Pro Pro Met Gln Arg Glu Leu Arg Ser Phe 385 390 395 400

Tyr Gly Gln Ala Pro Ala Val Val Glu Ile Cys Ala Gln Ala Gly Ala 405 410 415

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Pro Ser Ser Val Lys Glu Ala Glu Met Val Val
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<211> 540

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<213> Arabidopsis thaliana

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acg Thr	agg Arg 50	cgg Arg	ttc Phe	ctg Leu	cac His	gaa Glu 55	ggt Gly	cca Pro	gat Asp	acc Thr	gtg Val 60	gag Glu	gag Glu	ctt Leu	ctc Leu	192
gaa Glu 65	aga Arg	cat His	cta Leu	gcg Ala	aag Lys 70	aaa Lys	gag Glu	aaa Lys	cca Pro	ata Ile 75	atc Ile	gat Asp	cac His	gat Asp	gag Glu 80	240
gct Ala	gag Glu	ttt Phe	ctg Leu	aat Asn 85	aga Arg	cgg Arg	cgt Arg	ctg Leu	acg Thr 90	agc Ser	acg Thr	cgc Arg	cgg Arg	gaa Glu 95	gcg Ala	288
ttg Leu	agt Ser	ttg Leu	tac Tyr 100	aga Arg	gac Asp	ata Ile	tta Leu	cga Arg 105	gcg Ala	act Thr	cgg Arg	ttc Phe	ttc Phe 110	acg Thr	tgg Trp	336
att Ile	gat Asp	tcc Ser 115	agg Arg	gga Gly	aat Asn	tta Leu	tgg Trp 120	agg Arg	gac Asp	gtg Val	ttg Leu	aga Arg 125	gag Glu	aac Asn	gcg Ala	384
agg Arg	aag Lys 130	gag Glu	ttt Phe	gaa Glu	gcg Ala	gcg Ala 135	cga Arg	ttt Phe	gag Glu	acg Thr	gat Asp 140	ccg Pro	gag Glu	gtt Val	atc Ile	432
aca Thr 145	agg Arg	ctt Leu	ctg Leu	ata Ile	ggt Gly 150	gga Gly	agc Ser	gac Asp	gcc Ala	gtt Val 155	tcg Ser	tct Ser	gct Ala	tta Leu	gat Asp 160	480
aag Lys	ctt Leu	gcg Ala	gag Glu	aag Lys 165	caa Gln	aga Arg	gag Glu	atg Met	att Ile 170	gag Glu	aaa Lys	caa Gln	cgc Arg	cgt Arg 175	ggt Gly	528
_	caa Gln	_	tga													540
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<212> PRT

<213> Arabidopsis thaliana

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Cys Cys Cys Leu Arg Glu Met Met Ala Ala Lys Leu Gln Lys Trp Arg 20 25 30

Asn Leu Ala Gly Arg Leu Asp Leu Met Asn Arg Ser Gly Ala Val Ser 35 40 45

Thr Arg Arg Phe Leu His Glu Gly Pro Asp Thr Val Glu Glu Leu Leu 50 55 60

Glu Arg His Leu Ala Lys Lys Glu Lys Pro Ile Ile Asp His Asp Glu 65 70 75 80

Ala Glu Phe Leu Asn Arg Arg Arg Leu Thr Ser Thr Arg Arg Glu Ala 85 90 95

Leu Ser Leu Tyr Arg Asp Ile Leu Arg Ala Thr Arg Phe Phe Thr Trp 100 105 110

Ile Asp Ser Arg Gly Asn Leu Trp Arg Asp Val Leu Arg Glu Asn Ala 115 120 125

Arg Lys Glu Phe Glu Ala Ala Arg Phe Glu Thr Asp Pro Glu Val Ile 130 135 140

Thr Arg Leu Leu Ile Gly Gly Ser Asp Ala Val Ser Ser Ala Leu Asp 145 150 155 160

Lys Leu Ala Glu Lys Gln Arg Glu Met Ile Glu Lys Gln Arg Gly 165 170 175

Asp Gln Arg

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<213> Arabidopsis thaliana

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gct Ala	cat His	cag Gln 35	att Ile	cca Pro	gat Asp	acc Thr	ctc Leu 40	ctc Leu	tcg Ser	ctt Leu	cag Gln	cat His 45	cca Pro	cca Pro	act Thr	144	4
tat Tyr	acg Thr 50	ctc Leu	gga Gly	aag Lys	cgt Arg	aga Arg 55	acc Thr	gat Asp	cac His	aat Asn	cta Leu 60	ctt Leu	atc Ile	cct Pro	gaa Glu	192	2
tct Ser 65	gaa Glu	ctt Leu	aca Thr	aaa Lys	atc Ile 70	gga Gly	gct Ala	gaa Glu	ctt Leu	cat His 75	tat Tyr	act Thr	caa Gln	aga Arg	gga Gly 80	24	0
gga Gly	gac Asp	atc Ile	acc Thr	ttc Phe 85	cat His	ggc Gly	cct Pro	cat His	caa Gln 90	gcc Ala	atc Ile	tta Leu	tat Tyr	ccc Pro 95	atc Ile	28	8
att Ile	tcc Ser	tta Leu	cgc Arg 100	agc Ser	att Ile	ggt Gly	ttt Phe	ggt Gly 105	gct Ala	agg Arg	aac Asn	tac Tyr	gtg Val 110	gag Glu	aca Thr	33	6
ttg Leu	gag Glu	cgg Arg 115	tca Ser	atg Met	atc Ile	gag Glu	ttt Phe 120	gct Ala	tcg Ser	att Ile	tat Tyr	ggc Gly 125	gtg Val	aaa Lys	gct Ala	38	4
cga Arg	gca Ala 130	gga Gly	aac Asn	aaa Lys	tgt Cys	gag Glu 135	act Thr	Gly aaa	gtt Val	tgg Trp	gtt Val 140	ggg	gat Asp	agg Arg	aag Lys	43	2
atc Ile 145	ggt Gly	gct Ala	att Ile	gly ggg	gtt Val 150	agg Arg	ata Ile	tct Ser	tct Ser	gga Gly 155	atc Ile	act Thr	agt Ser	cat His	ggt Gly 160	48	0
ttg Leu	gcc Ala	tta Leu	aat Asn	ata Ile 165	gat Asp	cct Pro	gat Asp	atg Met	aag Lys 170	tac Tyr	ttt Phe	gag Glu	cac His	att Ile 175	gtg Val	52	8
cct Pro	tgt Cys	Gly	att Ile 180	gct Ala	gat Asp	aaa Lys	gaa Glu	gtt Val 185	aca Thr	tct Ser	ttg Leu	cga Arg	aga Arg 190	gag Glu	acg Thr	57	6
gat Asp	act Thr	ctg Leu 195	Leu	cct Pro	tca Ser	gaa Glu	gaa Glu 200	gtg Val	att Ile	cat His	gaa Glu	cag Gln 205	Leu	gtt Val	tct Ser	62	4
tgt Cys	tta Leu 210	Ala	aaa Lys	gcg Ala	ttt Phe	tct Ser 215	Tyr	gat Asp	gat Asp	gtt Val	gtc Val 220	tgg Trp	aag Lys	gaa Glu	gat Asp	67	2
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<211> 235

<212> PRT

<213> Arabidopsis thaliana

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Tyr Leu Lys Ser Leu Lys Leu Gln Glu Lys Leu Val Ser Glu Arg Lys 20 25 30

Ala His Gln Ile Pro Asp Thr Leu Leu Ser Leu Gln His Pro Pro Thr 35 40 45

Tyr Thr Leu Gly Lys Arg Arg Thr Asp His Asn Leu Leu Ile Pro Glu 50 55 60

Ser Glu Leu Thr Lys Ile Gly Ala Glu Leu His Tyr Thr Gln Arg Gly 65 70 75 80

Gly Asp Ile Thr Phe His Gly Pro His Gln Ala Ile Leu Tyr Pro Ile 85 90 95

Ile Ser Leu Arg Ser Ile Gly Phe Gly Ala Arg Asn Tyr Val Glu Thr 100 105 110

Leu Glu Arg Ser Met Ile Glu Phe Ala Ser Ile Tyr Gly Val Lys Ala 115 120 125

Arg Ala Gly Asn Lys Cys Glu Thr Gly Val Trp Val Gly Asp Arg Lys 130 135 140

Ile Gly Ala Ile Gly Val Arg Ile Ser Ser Gly Ile Thr Ser His Gly 145 150 155 160

Leu Ala Leu Asn Ile Asp Pro Asp Met Lys Tyr Phe Glu His Ile Val 165 170 175

Pro Cys Gly Ile Ala Asp Lys Glu Val Thr Ser Leu Arg Arg Glu Thr 180 185 190

Asp Thr Leu Leu Pro Ser Glu Glu Val Ile His Glu Gln Leu Val Ser 195 200 205

Cys Leu Ala Lys Ala Phe Ser Tyr Asp Asp Val Val Trp Lys Glu Asp 210 215 220

Pro Ser Leu Ile Leu Asp Thr Gln Asp Lys Glu 225 230 235

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acg Thr	agt Ser	gat Asp 115	gat Asp	gat Asp	aat Asn	gat Asp	tct Ser 120	tca Ser	aag Lys	act Thr	ggt Gly	gtt Val 125	gaa Glu	ttg Leu	ctt Leu	384
tgt Cys	gtg Val 130	gtg Val	aga Arg	gct Ala	gtg Val	ttg Leu 135	aag Lys	aaa Lys	ata Ile	cga Arg	agg Arg 140	aga Arg	gtt Val	tta Leu	gtt Val	432
													aga Arg			480
atg Met	att Ile	gag Glu	aat Asn	gtg Val 165	ttt Phe	cat His	cga Arg	cgt Arg	tcg Ser 170	gag Glu	att Ile	ttg Leu	gat Asp	cca Pro 175	cct Pro	528
gtt Val	gcg Ala	aac Asn	gtt Val 180	gat Asp	cat His	ttg Leu	ctt Leu	gtt Val 185	ctt Leu	ttc Phe	tct Ser	ttg Leu	gat Asp 190	caa Gln	ccg Pro	576
aaa Lys	ctt Leu	gag Glu 195	ccg Pro	ttt Phe	act Thr	ctt Leu	act Thr 200	agg Arg	ttc Phe	ttg Leu	gtg Val	gaa Glu 205	gct Ala	gaa Glu	tct Ser	624
act Thr	cgt Arg 210	att Ile	cca Pro	tta Leu	aca Thr	ctt Leu 215	gct Ala	ttg Leu	aat Asn	aaa Lys	act Thr 220	gaa Glu	ctc Leu	att Ile	agt Ser	672
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gaa Glu	cca Pro	ttg Leu	ttt Phe	tgt Cys 245	agt Ser	gtg Val	gga Gly	act Thr	aaa Lys 250	gat Asp	gga Gly	ctt Leu	gat Asp	gat Asp 255	att Ile	768
gcg Ala	ttt Phe	gtt Val	ctg Leu 260	aga Arg	gat Asp	cag Gln	act Thr	tct Ser 265	gtg Val	att Ile	gtt Val	gga Gly	cct Pro 270	agt Ser	ggt Gly	816
gtt Val	gga Gly	aag Lys 275	tcg Ser	agt Ser	tta Leu	atc Ile	aac Asn 280	gta Val	ttg Leu	agg Arg	agt Ser	aat Asn 285	cat His	ggt Gly	ggt Gly	864
ggt Gly	gtg Val 290	gtg Val	gaa Glu	gat Asp	gag Glu	aat Asn 295	tgg Trp	ttt Phe	gag Glu	cct Pro	atg Met 300	tta Leu	ggt Gly	aat Asn	aag Lys	912
tgg Trp 305	ttt Phe	gat Asp	gat Asp	cag Gln	cga Arg 310	gta Val	gly aaa	gaa Glu	gtt Val	tcg Ser 315	agt Ser	aga Arg	agt Ser	ggt Gly	aga Arg 320	960
ggt Gly	aaa Lys	cat His	aca Thr	aca Thr 325	cga Arg	aat Asn	gta Val	tcg Ser	cta Leu 330	ctg Leu	ccg Pro	gtt Val	tct Ser	gaa Glu 335	ggt Gly	1008
ggt Gly	tac Tyr	ctc Leu	gct Ala 340	gat Asp	act Thr	cct Pro	ggc	ttt Phe 345	aac Asn	cag Gln	cct Pro	agt Ser	ttg Leu 350	ctg Leu	aaa Lys	1056

gta Val	acg Thr	aag Lys 355	cat His	tca Ser	cta Leu	gct Ala	cac His 360	tgt Cys	ttt Phe	cct Pro	gag Glu	ata Ile 365	cgg Arg	aac Asn	atg Met	1104
att Ile	gag Glu 370	agc Ser	gaa Glu	aaa Lys	tgt Cys	gga Gly 375	ttc Phe	aga Arg	gac Asp	tgc Cys	ttg Leu 380	cat His	att Ile	Gly aaa	gaa Glu	1152
cca Pro 385	gga Gly	tgt Cys	gtt Val	gtg Val	aaa Lys 390	ggt Gly	gac Asp	tgg Trp	gaa Glu	agg Arg 395	tat Tyr	cct Pro	tac Tyr	tac Tyr	tta Leu 400	1200
caa Gln	ttg Leu	ctt Leu	gat Asp	gag Glu 405	atc Ile	aga Arg	atc Ile	agg Arg	gaa Glu 410	gaa Glu	ttt Phe	cag Gln	ctt Leu	agg Arg 415	act Thr	1248
ttt Phe	gga Gly	acc Thr	aaa Lys 420	agg Arg	gaa Glu	gat Asp	gat Asp	gtt Val 425	agg Arg	tac Tyr	aag Lys	gtg Val	gga Gly 430	gac Asp	atg Met	1296
ggt Gly	gtg Val	aaa Lys 435	cat His	gct Ala	gaa Glu	cca Pro	cgg Arg 440	tta Leu	atg Met	cct Pro	aag Lys	aag Lys 445	cat His	agg Arg	aga Arg	1344
gag Glu	tca Ser 450	agg Arg	aag Lys	aaa Lys	acg Thr	aaa Lys 455	cag Gln	aca Thr	atg Met	atc Ile	agt Ser 460	gag Glu	ctg Leu	gat Asp	gag Glu	1392
ttc Phe 465	gaa Glu	gat Asp	gaa Glu	gac Asp	agt Ser 470	gat Asp	ttg Leu	tac Tyr	ata Ile	gag Glu 475	aac Asn	gac Asp	cca Pro	atc Ile	gtc Val 480	1440
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<211> 490

<212> PRT

<213> Arabidopsis thaliana

<400> 54

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Arg His Thr Ala Ile Phe His Gly Gly Val Gly Val Arg Phe Lys Phe 20 25 30

Leu Arg Ser Phe Ser Pro Leu Ser Ala Arg Arg Asp Asn Pro Asp Val 35 40 45

Ser Arg Lys Pro Gln Pro Ser Lys Asn Met Leu Arg Ala Lys His Ile Gly Lys Asn Tyr Ser Ser Ser Leu Ser Pro Val Leu Ser Pro Glu His 70 Lys Pro Ser Leu Leu Glu Ser Gln Ala Ile Gly Thr Val Ala Thr Ala 90 85 Gln Ala Asn Phe Met Arg Val Ile Val Gln Asp Val Ala Asn Ser Val 105 100 Thr Ser Asp Asp Asp Asn Asp Ser Ser Lys Thr Gly Val Glu Leu Leu Cys Val Val Arg Ala Val Leu Lys Lys Ile Arg Arg Val Leu Val , 135 Gly Asp Lys Val Leu Val Gly Ser Ile Asp Trp Val Asp Arg Arg Gly 155 160 150 145 Met Ile Glu Asn Val Phe His Arg Arg Ser Glu Ile Leu Asp Pro Pro 170 165 Val Ala Asn Val Asp His Leu Leu Val Leu Phe Ser Leu Asp Gln Pro 190 185 180 Lys Leu Glu Pro Phe Thr Leu Thr Arg Phe Leu Val Glu Ala Glu Ser 200 Thr Arg Ile Pro Leu Thr Leu Ala Leu Asn Lys Thr Glu Leu Ile Ser 215 210 Glu Glu Glu Leu Glu Thr Trp Lys Ile Arg Leu Arg Gly Trp Asn Tyr 235 230 225 Glu Pro Leu Phe Cys Ser Val Gly Thr Lys Asp Gly Leu Asp Asp Ile 245 250 Ala Phe Val Leu Arg Asp Gln Thr Ser Val Ile Val Gly Pro Ser Gly 260 270 Val Gly Lys Ser Ser Leu Ile Asn Val Leu Arg Ser Asn His Gly Gly 280

275

Gly Val Val Glu Asp Glu Asn Trp Phe Glu Pro Met Leu Gly Asn Lys 290 295 300

Trp Phe Asp Asp Gln Arg Val Gly Glu Val Ser Ser Arg Ser Gly Arg 305 310 315 320

Gly Lys His Thr Thr Arg Asn Val Ser Leu Leu Pro Val Ser Glu Gly 325 330 335

Gly Tyr Leu Ala Asp Thr Pro Gly Phe Asn Gln Pro Ser Leu Leu Lys 340 345 350

Val Thr Lys His Ser Leu Ala His Cys Phe Pro Glu Ile Arg Asn Met 355 360 365

Ile Glu Ser Glu Lys Cys Gly Phe Arg Asp Cys Leu His Ile Gly Glu 370 375 380

Pro Gly Cys Val Val Lys Gly Asp Trp Glu Arg Tyr Pro Tyr Tyr Leu 385 390 395 400

Gln Leu Leu Asp Glu Ile Arg Ile Arg Glu Glu Phe Gln Leu Arg Thr 405 410 415

Phe Gly Thr Lys Arg Glu Asp Asp Val Arg Tyr Lys Val Gly Asp Met 420 425 430

Gly Val Lys His Ala Glu Pro Arg Leu Met Pro Lys Lys His Arg Arg 435 440 445

Glu Ser Arg Lys Lys Thr Lys Gln Thr Met Ile Ser Glu Leu Asp Glu 450 455 460

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gtt Val	gct Ala	att Ile 35	gtt Val	aat Asn	ctg Leu	gat Asp	cct Pro 40	gca Ala	aat Asn	gat Asp	gca Ala	tta Leu 45	cct Pro	tat Tyr	gag Glu	1	44
tgt Cys	ggt Gly 50	gtg Val	aat Asn	ata Ile	gaa Glu	gaa Glu 55	ttg Leu	atc Ile	aag Lys	tta Leu	gaa Glu 60	gat Asp	gtt Val	atg Met	tcg Ser	1	92
gaa Glu 65	cac His	tcg Ser	ctt Leu	ggt Gly	cct Pro 70	aat Asn	gga Gly	ggt Gly	ctt Leu	gta Val 75	tat Tyr	tgt Cys	atg Met	gag Glu	tac Tyr 80	2	40
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aag Lys	gat Asp	cat His	tac Tyr 100	att Ile	ctc Leu	ttt Phe	gat Asp	ttt Phe 105	cct Pro	ggc Gly	caa Gln	gtg Val	gaa Glu 110	ttg Leu	ttc Phe	3	36
ttc Phe	att Ile	cat His 115	gac Asp	agt Ser	acc Thr	aag Lys	aat Asn 120	gtt Val	ctc Leu	acg Thr	aag Lys	ctg Leu 125	att Ile	aaa Lys	tca Ser	3	84
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tgt Cys 145	gat Asp	ccc Pro	Gly ggg	aac Asn	tac Tyr 150	gta Val	agt Ser	tcg Ser	cta Leu	ctt Leu 155	ctc Leu	tcc Ser	tta Leu	tcc Ser	aca Thr 160	4	80
atg Met	ctt Leu	cac His	atg Met	gaa Glu 165	ctc Leu	cca Pro	cat His	gtc Val	aat Asn 170	gta Val	ttg Leu	tct Ser	aaa Lys	atc Ile 175	gat Asp	5	28

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gat gtt Asp Val	caa Gln 195	gac Asp	ttg Leu	tca Ser	tac Tyr	ttg Leu 200	gag Glu	cac His	cat His	ctt Leu	agt Ser 205	caa Gln	gat Asp	cct Pro	624
cgc tct Arg Ser 210	Ala	aag Lys	tac Tyr	aga Arg	aaa Lys 215	cta Leu	aca Thr	aaa Lys	gag Glu	cta Leu 220	tgt Cys	agt Ser	gtc Val	att Ile	672
gaa gat Glu Asj 225	tac Tyr	agt Ser	ctt Leu	gtt Val 230	aat Asn	ttt Phe	aca Thr	acc Thr	ttg Leu 235	gat Asp	att Ile	cag Gln	gat Asp	aaa Lys 240	720
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ata tt	gcc Ala	ggc Gly 260	att Ile	gat Asp	gca Ala	agt Ser	gtg Val 265	gtt Val	gaa Glu	tac Tyr	agc Ser	aag Lys 270	att Ile	gca Ala	816
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Glu His Ser Leu Gly Pro Asn Gly Gly Leu Val Tyr Cys Met Glu Tyr Leu Glu Lys Asn Ile Asp Trp Leu Glu Ser Lys Leu Lys Pro Leu Leu Lys Asp His Tyr Ile Leu Phe Asp Phe Pro Gly Gln Val Glu Leu Phe Phe Ile His Asp Ser Thr Lys Asn Val Leu Thr Lys Leu Ile Lys Ser Leu Asn Leu Arg Leu Thr Ala Val Gln Leu Ile Asp Ser His Leu Cys Cys Asp Pro Gly Asn Tyr Val Ser Ser Leu Leu Leu Ser Leu Ser Thr Met Leu His Met Glu Leu Pro His Val Asn Val Leu Ser Lys Ile Asp Leu Ile Gly Ser Tyr Gly Lys Leu Ala Phe Asn Leu Asp Phe Tyr Thr Asp Val Gln Asp Leu Ser Tyr Leu Glu His His Leu Ser Gln Asp Pro Arg Ser Ala Lys Tyr Arg Lys Leu Thr Lys Glu Leu Cys Ser Val Ile Glu Asp Tyr Ser Leu Val Asn Phe Thr Thr Leu Asp Ile Gln Asp Lys Glu Ser Val Gly Asp Leu Val Lys Leu Ile Asp Lys Ser Asn Gly Tyr Ile Phe Ala Gly Ile Asp Ala Ser Val Val Glu Tyr Ser Lys Ile Ala Ile Gly Gln Thr Asp Trp Asp Tyr Asn Arg Val Ala Ala Val Gln Glu

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Asp Leu Val Glu Thr Asp Met Val Ser Pro Asp Ile Ser Lys His His 145 150 155 160													
aaa gca aag gag cct ctc ttg gtt tct cag cca caa tgc tgc aga aca Lys Ala Lys Glu Pro Leu Leu Val Ser Gln Pro Gln Cys Cys Arg Thr 165 170 175	528												
acc tac gat gga tca agt agt tct gct agt tgt aca ttt caa gct ctt Thr Tyr Asp Gly Ser Ser Ser Ser Ala Ser Cys Thr Phe Gln Ala Leu 180 185 190	576												
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cgc gcc tgc att gta gcc tca cac cca aca acc ggt tta tcc ttc agc Arg Ala Cys Ile Val Ala Ser His Pro Thr Thr Gly Leu Ser Phe Ser 210 215 220	672												
cta act ttt ata aat aac cca aat ggt gaa gaa tct gag ctg ctt tac Leu Thr Phe Ile Asn Asn Pro Asn Gly Glu Glu Ser Glu Leu Leu Tyr 225 230 235 240	720												
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Arg Gly Glu Asn Gln Ser Ser Leu Glu Met Val Glu Lys Leu Arg Asn Glu Ile Ile Ser Ile Arg Ser Gly Arg Asp Asp Lys Phe Leu Glu Cys Gln Lys Leu Leu Met Glu Glu Glu Leu Lys Asn Lys Ser Leu Ser Glu Glu Val Val Lys Leu Lys Glu Leu Val Glu Glu His Pro Arg Asn Tyr Glu Asp Gln Ser Gly Lys Lys Gln Lys Arg Lys Thr Pro Glu Ser Ala Arg Val Thr Thr Arg Ser Met Ile Lys Arg Ser Arg Leu Ser Glu Asp Leu Val Glu Thr Asp Met Val Ser Pro Asp Ile Ser Lys His His Lys Ala Lys Glu Pro Leu Leu Val Ser Gln Pro Gln Cys Cys Arg Thr Thr Tyr Asp Gly Ser Ser Ser Ser Ala Ser Cys Thr Phe Gln Ala Leu Gly Lys His Leu Leu Gly Met Lys Leu Ser Thr Asn Asn Lys Gly Lys Arg Ala Cys Ile Val Ala Ser His Pro Thr Thr Gly Leu Ser Phe Ser Leu Thr Phe Ile Asn Asn Pro Asn Gly Glu Glu Ser Glu Leu Leu Tyr Lys Pro Ala Ser Leu Gly Thr Phe Gln Arg Val Ala Pro Glu Trp Met Arg Glu Val Ile Lys Phe Ser Thr Ser Met Cys Pro Ile Phe Phe Glu Arg Val Ser Arg Val Ile Lys Leu Asn Cys

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<222> (1)..(1467)

<223> 37351

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gtg Val	gcg Ala	aca Thr	ctt Leu 20	gtt Val	ata Ile	gcc Ala	aaa Lys	ctc Leu 25	atc Ile	ttc Phe	tct Ser	ttc Phe	ttc Phe 30	act Thr	tct Ser		96
gat Asp	tct Ser	aag Lys 35	aag Lys	aag Lys	cgt Arg	ctt Leu	cct Pro 40	cct Pro	act Thr	ctt Leu	aaa Lys	gct Ala 45	tgg Trp	cct Pro	cca Pro	;	144
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cac	aaa Lys	aag Lys	att Ile	act Thr 85	ttt Phe	ctt Leu	att Ile	ggt Gly	cct Pro 90	gaa Glu	gtc Val	tct Ser	gct Ala	cat His 95	ttt Phe	:	288
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gtt Val	cgt Arg 130	cag Gln	gag Glu	cag Gln	ttt Phe	cgg Arg 135	ttc Phe	ttc Phe	act Thr	gag Glu	gca Ala 140	ctt Leu	aga Arg	gtt Val	aac Asn		432
aag Lys 145	ttg Leu	aag Lys	ggt Gly	tat Tyr	gtg Val 150	gat Asp	atg Met	atg Met	gtt Val	act Thr 155	gaa Glu	gct Ala	gag Glu	gat Asp	tac Tyr 160		480

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ttc Phe	gca Ala	aaa Lys	atc Ile	att Ile 245	gly ggg	tcg Ser	aga Arg	aaa Lys	cgc Arg 250	tct Ser	gga Gly	aaa Lys	aca Thr	gag Glu 255	aac Asn	768
gac Asp	atg Met	ctg Leu	cag Gln 260	tgt Cys	ttc Phe	atc Ile	gaa Glu	tca Ser 265	aag Lys	tac Tyr	aaa Lys	gat Asp	ggt Gly 270	aga Arg	cag Gln	816
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acc Thr 385	tcc Ser	cct Pro	gca Ala	ttt Phe	gcc Ala 390	aac Asn	cgc Arg	tta Leu	ccg Pro	cac His 395	atc Ile	ttc Phe	aaa Lys	gac Asp	ccc Pro 400	1200

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gcc Ala	gca Ala	gly aaa	gca Ala 420	ttc Phe	tcg Ser	tac Tyr	att Ile	gca Ala 425	ttc Phe	gga Gly	gly ggg	gga Gly	agg Arg 430	cac His	gly aaa	1296
tgc Cys	ctt Leu	gga Gly 435	gag Glu	ccg Pro	ttt Phe	gct Ala	tac Tyr 440	ctg Leu	cag Gln	atc Ile	aaa Lys	gcc Ala 445	ata Ile	tgg Trp	agt Ser	1344
cat His	ttg Leu 450	ttg Leu	agg Arg	aac Asn	ttc Phe	gag Glu 455	ctt Leu	gag Glu	cta Leu	gtt Val	tca Ser 460	ccg Pro	ttc Phe	cct Pro	gag Glu	1392
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Asp Ser Lys Lys Lys Arg Leu Pro Pro Thr Leu Lys Ala Trp Pro Pro 35 40 45

Leu Val Gly Ser Leu Ile Lys Phe Leu Lys Gly Pro Ile Ile Met Leu 50 55 60

Arg Glu Glu Tyr Pro Lys Leu Gly Ser Val Phe Thr Val Asn Leu Val 65 70 75 80

His Lys Lys Ile Thr Phe Leu Ile Gly Pro Glu Val Ser Ala His Phe 85 90 95

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Val	Arg 130	Gln	Glu	Gln	Phe	Arg 135	Phe	Phe	Thr	Glu	Ala 140	Leu	Arg	Val	Asn
Lys 145	Leu	Lys	Gly	Tyr	Val 150	Asp	Met	Met	Val	Thr 155	Glu	Ala	Glu	Asp	Tyr 160
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Val	Arg	Asp 195	Gln	Leu	Phe	Asp	Asp 200	Val	Ser	Ala	Leu	Phe 205	His	Asp	Leu
Asp	Asn 210	Gly	Met	Leu	Pro	Ile 215	Ser	Val	Leu	Phe	Pro 220	Tyr	Leu	Pro	Ile
Pro 225	Ala	His	Arg	Arg	Arg 230	Asp	Arg	Ala	Arg	Glu 235	Lys	Leu	Ser	Glu	Ile 240
Phe	Ala	Lys	Ile	Ile 245	Gly	Ser	Arg	Lys	Arg 250	Ser	Gly	Lys	Thr	Glu 255	Asn
Asp	Met	Leu	Gln 260	Cys	Phe	Ile	Glu	Ser 265	Lys	Tyr	Lys	Авр	Gly 270	Arg	Gln
Thr	Thr	Glu 275	Ser	Glu	Val	Thr	Gly 280	Leu	Leu	Ile	Ala	Ala 285	Leu	Phe	Ala
Gly	Gln 290	His	Thr	Ser	Ser	Ile 295	Thr	Ser	Thr	Trp	Thr 300	Gly	Ala	Tyr	Leu
Met 305	Arg	Tyr	Lys	Glu	Tyr 310	Phe	Ser	Ala	Ala	Leu 315	Asp	Glu	Gln	Lys	Asn 320
Leu	Ile	Ala	Lys	His	Gly	Asp	Lys	Ile	Asp	His	Asp	Ile	Leu	Ser	Glu

Met Asp Val Leu Tyr Arg Cys Ile Lys Glu Ala Leu Arg Leu His Pro 340 345 350

Pro Leu Ile Met Leu Met Arg Ala Ser His Ser Asp Phe Ser Val Thr 355 360 365

Ala Arg Asp Gly Lys Thr Tyr Asp Ile Pro Lys Gly His Ile Val Ala 370 380

Thr Ser Pro Ala Phe Ala Asn Arg Leu Pro His Ile Phe Lys Asp Pro 385 395 400

Asp Thr Tyr Asp Pro Glu Arg Phe Ser Pro Gly Arg Glu Glu Asp Lys 405 410 415

Ala Ala Gly Ala Phe Ser Tyr Ile Ala Phe Gly Gly Gly Arg His Gly
420 425 430

Cys Leu Gly Glu Pro Phe Ala Tyr Leu Gln Ile Lys Ala Ile Trp Ser 435 440 445

His Leu Leu Arg Asn Phe Glu Leu Glu Leu Val Ser Pro Phe Pro Glu 450 455 460

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Arg Tyr Lys Arg Arg Gln Leu Ser 485

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<223> 37389

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cca Pro																192
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agt Ser	gat Asp	gat Asp	gag Glu	gaa Glu 85	gaa Glu	gaa Glu	gaa Glu	gat Asp	cat His 90	agt Ser	caa Gln	atc Ile	tgt Cys	aca Thr 95	gcg Ala	288
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gct Ala	gag Glu 130	agt Ser	gcc Ala	cta Leu	tat Tyr	gag Glu 135	gtt Val	atc Ile	aac Asn	gac Asp	cac His 140	caa Gln	acc Thr	gaa Glu	atc Ile	432
aag Lys 145	gac Asp	gac Asp	att Ile	agg Arg	aat Asn 150	caa Gln	gta Val	tca Ser	gtt Val	gtt Val 155	gaa Glu	aca Thr	gaa Glu	ata Ile	atg Met 160	480
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cga Arg	aaa Lys	gtt Val 195	gct Ala	gaa Glu	gca Ala	ctt Leu	gat Asp 200	acc Thr	cat His	ctg Leu	act Thr	gca Ala 205	gtc Val	caa Gln	cgc Arg	624
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ctt Leu	gcg Ala 290	gaa Glu	cag Gln	aag Lys	gct Ala	gtg Val 295	ata Ile	gag Glu	agt Ser	gtt Val	acg Thr 300	gly aaa	agt Ser	tca Ser	gct Ala	912
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gaa Glu	ttc Phe	cac His 435	Lys	gct Ala	tgc Cys	att Ile	tac Tyr 440	Thr	gtc Val	cca Pro	aag Lys	cat His 445	тте	gta Val	aac Asn	1344
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cag ttt ctg Gln Phe Leu 530	aag gtt gtg Lys Val Va	g aat gtt gt . Asn Val Va 535	ıl Arg Glu H	eat ttc ttg His Phe Leu 540	cag aaa 1632 Gln Lys
ttg cgg gcg Leu Arg Ala 545	aag aag gad Lys Lys Asp 550	Thr Ser As	it cta ctt g sp Leu Leu V 555	gtg atc ata Val Ile Ile	gcc gaa 1680 Ala Glu 560
atc aca gcg Ile Thr Ala	tac tta gat Tyr Leu Asp 565	gac cgg at Asp Arg Me	g tat ctc a t Tyr Leu L 570	aag gaa cct Lys Glu Pro	gaa gga 1728 Glu Gly 575
aga gct atg Arg Ala Met	aag acg act Lys Thr Thi 580	agt acc tt Ser Thr Le 58	eu Ser Ser G	gaa ctt act Blu Leu Thr 590	gct gaa 1776 Ala Glu
tta aat cag Leu Asn Gln 595	ccg aac tac Pro Asn Ty	e aat cag aa Asn Gln As 600	at tac cag a sn Tyr Gln A	agg aat gat Arg Asn Asp 605	tac aga 1824 Tyr Arg
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Ser Ile Asp	Pro Glu Pro 20	Asn Trp As 25		Ser Leu Val 30	Ala Glu

Ile Ala Ser Val Glu Lys Lys Leu Asn Gly Phe Ser Met Tyr Pro Gln 35 40 45

- Pro Ile Thr Asn Thr Thr Leu Arg Met Gly Arg Arg Gly Gly Phe 50 55 60
- Val Met His Val Ser Glu Asp Glu Met Glu Ser Asp Glu Gly Glu Glu 65 70 75 80
- Ser Asp Asp Glu Glu Glu Glu Glu Asp His Ser Gln Ile Cys Thr Ala 85 90 95
- Gly Lys Arg Phe Ala Cys Asp Glu Leu Tyr Leu Ser Asp Glu Ser Asp 100 105 110
- Glu Glu Phe Asp His Glu Pro Glu Tyr Met Met Asn Lys Leu Gly Leu 115 120 125
- Ala Glu Ser Ala Leu Tyr Glu Val Ile Asn Asp His Gln Thr Glu Ile 130 135 140
- Lys Asp Asp Ile Arg Asn Gln Val Ser Val Val Glu Thr Glu Ile Met 145 150 155 160
- Asn Glu Ile Glu Thr Ser Leu Ser Ala Ile Ala Arg Val Glu Lys Tyr 165 170 175
- Ser Glu Thr Arg Lys Glu Val Glu Arg Lys Leu Asp Leu Gln Tyr Gln 180 185 190
- Arg Lys Val Ala Glu Ala Leu Asp Thr His Leu Thr Ala Val Gln Arg 195 200 205
- Glu His Lys Ile Lys Ser Gln Ile Glu Glu Arg Lys Ile Arg Ser Glu 210 215 220
- Glu Ala Gln Glu Glu Ala Arg Arg Lys Glu Arg Ala His Gln Glu Glu 225 230 235 240
- Lys Ile Arg Gln Glu Lys Ala Arg Ala Glu Ala Gln Met Leu Ala Lys 245 250 255
- Ile Arg Ala Glu Glu Glu Lys Lys Glu Val Glu Arg Lys Ala Ala Arg 260 265 270

Glu Val Ala Glu Lys Glu Val Ala Asp Arg Lys Ala Ala Glu Gln Lys 280 275 . Leu Ala Glu Gln Lys Ala Val Ile Glu Ser Val Thr Gly Ser Ser Ala 295 300 Thr Ser Asn Ala Gln Ala Gly Gly Asn Ser Ile Arg Ala Ala Glu Ser 315 310 Ala Leu Ile Leu Glu Asn His Arg Leu Lys Lys Leu Glu Glu Leu Glu 330 325 Thr Thr Asn Gln Ser Leu Lys Ser Arg Ser Asn Glu Asn Phe Ser Ser 345 Phe Glu Lys His Ile Gly Arg Val Ile Arg Gln Ile Ser Gly Thr Lys 355 Asp Ser Val Ser Gly Lys Ile Asn Asp Ile Val Lys Ile Phe Lys Asp 375 370 Pro Arg Cys Pro Val Ser Ile Ser Ile Ala Ala Phe Ala Lys Lys Met 400 395 390 385 Val Thr Thr Lys Glu Lys Pro Asn Pro Phe Ala Cys Ser Tyr Val Ile 410 415 405 Val Tyr Ile Asn Ser Gln Phe Pro Gln Val Met Asp Ile Leu Leu Ala 425 420 Glu Phe His Lys Ala Cys Ile Tyr Thr Val Pro Lys His Ile Val Asn 440 435 Ser Gln Ser Ala Trp Asp Ser Asp Ala Tyr Glu Arg Leu Asp Ser Ile 455 450

Met Arg Leu Tyr Gly Ala Leu Val Gln Thr Asp Ile Arg Val Gly Asn 465 470 475 480

Ala Thr Asn Val His Gly Ile Glu His Gly Trp Ala Trp Leu Ala Arg 485 490 495

Phe Leu Asn Lys Ile Pro Ala Asn Arg Ala Thr Ala Thr Ala Leu Asn 500 505 510

Ser Phe Leu Gln Thr Ala Gly Phe Gly Leu His Gln Arg Tyr Lys Ser 520 Gln Phe Leu Lys Val Val Asn Val Val Arg Glu His Phe Leu Gln Lys 535 Leu Arg Ala Lys Lys Asp Thr Ser Asp Leu Leu Val Ile Ile Ala Glu 555 545 550 Ile Thr Ala Tyr Leu Asp Asp Arg Met Tyr Leu Lys Glu Pro Glu Gly 570 565 Arg Ala Met Lys Thr Thr Ser Thr Leu Ser Ser Glu Leu Thr Ala Glu 585 Leu Asn Gln Pro Asn Tyr Asn Gln Asn Tyr Gln Arg Asn Asp Tyr Arg 600 Asn Tyr Tyr 610 <210> 63 <211> 1152 <212> DNA <213> Arabidopsis thaliana <220> <221> CDS <222> (1)..(1152) <223> 38108 <400> 63 atg gca acg gct tct cct cca ttt atc tca act ctc agc ttc act cac 48 Met Ala Thr Ala Ser Pro Pro Phe Ile Ser Thr Leu Ser Phe Thr His 10 5 tct tct ttc aaa act tct tct tct tct tca ttt tct ccg aag ctt ctt 96 Ser Ser Phe Lys Thr Ser Ser Ser Ser Phe Ser Pro Lys Leu Leu 20 cga ccc ctc tta agc ttt tcc gtc aaa gct tcc aga aag caa gta gag 144 Arg Pro Leu Leu Ser Phe Ser Val Lys Ala Ser Arg Lys Gln Val Glu 40 45 35

ata	gtg	ttt	gat	cct	gat	gag	agg	ctt	aat	aag	ata	ggt	gat	gat	gtt	192
Ile	Val 50	Phe	Asp	Pro	Asp	Glu 55	Arg	Leu	Asn	Lys	Ile 60	Gly	Asp	Asp	Val	
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atc Ile	aat Asn	gtt Val	ttc Phe	ttg Leu 85	agg Arg	atc Ile	acc Thr	gga Gly	aag Lys 90	cga Arg	gaa Glu	gat Asp	gly aaa	ttt Phe 95	cat His	288
gat Asp	tta Leu	gcc Ala	tct Ser 100	ttg Leu	ttt Phe	cat His	gtg Val	att Ile 105	agc Ser	tta Leu	gga Gly	gac Asp	act Thr 110	att Ile	aaa Lys	336
ttc Phe	tca Ser	ttg Leu 115	tca Ser	cca Pro	tca Ser	aag Lys	tct Ser 120	aaa Lys	gat Asp	cgt Arg	ttg Leu	tct Ser 125	act Thr	aac Asn	gtt Val	384
caa Gln	gga Gly 130	gtc Val	cct Pro	gtt Val	gat Asp	999 Gly 135	aga Arg	aat Asn	ctg Leu	att Ile	ata Ile 140	aaa Lys	gca Ala	ctt Leu	aac Asn	432
ctt Leu 145	tac Tyr	agg Arg	aag Lys	aaa Lys	act Thr 150	ggt Gly	agt Ser	aac Asn	aga Arg	ttc Phe 155	ttc Phe	tgg Trp	att Ile	cat His	tta Leu 160	480
gat Asp	aag Lys	aag Lys	gtg Val	cct Pro 165	acc Thr	gly ggg	gct Ala	gga Gly	ctc Leu 170	ggt Gly	ggt Gly	gga Gly	agt Ser	agt Ser 175	aat Asn	528
gct Ala	gca Ala	act Thr	gca Ala 180	ctc Leu	tgg Trp	gcg Ala	gca Ala	aat Asn 185	gag Glu	ctc Leu	aat Asn	gga Gly	ggt Gly 190	ctt Leu	gtc Val	576
act Thr	gag Glu	aac Asn 195	gaa Glu	ctc Leu	cag Gln	gat Asp	tgg Trp 200	tca Ser	agt Ser	gaa Glu	att Ile	999 Gly 205	tca Ser	gat Asp	att Ile	624
cct Pro	ttc Phe 210	ttc Phe	ttc Phe	tcg Ser	cat His	gga Gly 215	gct Ala	gcc Ala	tat Tyr	tgt Cys	acc Thr 220	Gjà aaa	aga Arg	ggt Gly	gag Glu	672
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ctc Leu	ata Ile	aag Lys	ccc Pro	cga Arg 245	gaa Glu	gca Ala	tgt Cys	tcc Ser	act Thr 250	gct Ala	gaa Glu	gtt Val	tac Tyr	aaa Lys 255	cgt Arg	768
ctt Leu	cgt Arg	tta Leu	gat Asp 260	cag Gln	acg Thr	agc Ser	aat Asn	att Ile 265	Asn	ccc Pro	ttg Leu	aca Thr	tta Leu 270	cta Leu	gag Glu	816
aat Asn	gtg Val	acc Thr 275	Ser	aat Asn	ggt Gly	gtg Val	tct Ser 280	Gln	agc Ser	ata Ile	tgc Cys	gta Val 285	aac Asn	gat Asp	ttg Leu	864

Glu F	_	_	g ttt .a Phe		_					_	_	_		912
		_	a tct a Ser		-		-	-	-			_		960
			c act r Thr 325							_				1008
		_	t gat p Asp 0	_	_		_			_		_	_	1056
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Ile Val Phe Asp Pro Asp Glu Arg Leu Asn Lys Ile Gly Asp Asp Val 50 55 60

Asp Lys Glu Ala Pro Leu Ser Arg Leu Lys Leu Phe Ser Pro Cys Lys 65 70 75 80

Ile Asn Val Phe Leu Arg Ile Thr Gly Lys Arg Glu Asp Gly Phe His
85 90 95

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Phe	Ser	Leu 115	Ser	Pro	Ser	Lys	Ser 120	Lys	Asp	Arg	Ļeu	Ser 125	Thr	Asn	Val
Gln	Gly 130	Val	Pro	Val	Asp	Gly 135	Arg	Asn	Leu	Ile	Ile 140	Lys	Ala	Leu	Asn
Leu 145	Tyr	Arg	Lys	Lys	Thr 150	Gly	Ser	Asn	Arg	Phe 155	Phe	Trp	Ile	His	Leu 160
Asp	Lys	Lys	Val	Pro 165	Thr	Gly	Ala	Gly	Leu 170	Gly	Gly	Gly	Ser	Ser 175	Asn
Ala	Ala	Thr	Ala 180	Leu	Trp	Ala	Ala	Asn 185	Glu	Leu	Asn	Gly	Gly 190	Leu	Val
Thr	Glu	Asn 195	Glu	Leu	Gln	Asp	Trp 200	Ser	Ser	Glu	Ile	Gly 205	Ser	Asp	Ile
Pro	Phe 210	Phe	Phe	Ser	His	Gly 215	Ala	Ala	Tyr	Cys	Thr 220	Gly	Arg	Gly	Glu
Ile 225	Val	Gln	Asp	Leu	Pro 230	Pro	Pro	Phe	Pro	Leu 235	Asp	Leu	Pro	Met	Val 240
Leu	Ile	Lys	Pro	Arg 245	Glu	Ala	Cys	Ser	Thr 250	Ala	Glu	Val	Tyr	Lys 255	Arg
Leu	Arg	Leu	Asp 260	Gln	Thr	Ser	Asn	Ile 265	Asn	Pro	Leu	Thr	Leu 270	Leu	Glu
Asn	Val	Thr 275	Ser	Asn	Gly	Val	Ser 280	Gln	Ser	Ile	Cys	Val 285	Asn	Asp	Leu
Glu	Pro 290	Pro	Ala	Phe	Ser	Val 295	Leu	Pro	Ser	Leu	Tys	Arg	Leu	Lys	Gln
Arg 305	Ile	Ile	Ala	Ser	Gly 310	Arg	Gly	Glu	Tyr	Asp 315	Ala	Val	Phe	Met	Ser 320
Gly	Ser	Gly	Ser	Thr 325	Ile	Ile	Gly	Ile	Gly 330	Ser	Pro	Asp	Pro	Pro	Gln

Phe Ile Tyr Asp Asp Glu Glu Tyr Lys Asn Val Phe Leu Ser Glu Ala 340 345 350

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336

cca act ttc att gta cgg aaa aga cca gta aag ctc agt tct ctt aac

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		Gly				_		_		_		Leu	_		cga Arg	432
	att Ile															480
	gaa Glu															528
	gaa Glu															576
	caa Gln															624
	tgg Trp 210															672
	gat Asp															720
	aca Thr															768
	tcg Ser															816
	att Ile															864
tcg Ser	tct Ser 290	gga Gly	atc Ile	act Thr	cag Gln	gat Asp 295	ggt Gly	gaa Glu	gct Ala	ata Ile	cca Pro 300	aga Arg	ctt Leu	gca Ala	Gly aaa	912
	aga Arg															960
	atc Ile							Pro								1008
cgt	gtc	ctc	tcg	gat	aca	ttt	cct	cat	cac	tcg	gtt	gtg	gga	atc	gag	1056

Arg Val Leu Ser Asp Thr Phe Pro His His Ser Val Val Gly Ile Glu 340 345 350

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Asn Ala Arg Glu Ile Val Leu Ala Gly Gly Asn Ile His Cys Ile Thr
355
360
365

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Asp Val Ala Lys Ala Ile Ser Lys Phe Glu Pro Val Thr Val Cys Ala 50 55 60

Ser Pro Ala Gln Trp Glu Asn Ala Arg Lys Gln Leu Pro Glu Asp Ile 65 70 75 80

Arg Val Val Glu Met Ser Met Asn Asp Ser Trp Phe Arg Asp Ser Gly 85 90 95

Pro Thr Phe Ile Val Arg Lys Arg Pro Val Lys Leu Ser Ser Leu Asn 100 105 110

Arg Asn Ile Ala Gly Ile Asp Trp Asn Phe Asn Ala Trp Gly Gly Ala 115 120 125

Asn Asp Gly Cys Tyr Asn Asp Trp Ser His Asp Leu Leu Val Ser Arg 130 135 140

Lys Ile Leu Ala Leu Glu Arg Ile Pro Arg Phe Gln His Ser Met Ile Leu Glu Gly Gly Ser Ile His Val Asp Gly Glu Gly Thr Cys Leu Val Thr Glu Glu Cys Leu Leu Asn Lys Asn Arg Asn Pro His Met Ser Lys Glu Gln Ile Glu Glu Leu Lys Lys Tyr Leu Gly Val Gln Ser Phe Ile Trp Leu Pro Arg Gly Leu Tyr Gly Asp Glu Asp Thr Asn Gly His Ile Asp Asn Met Cys Cys Phe Ala Arg Pro Gly Val Val Leu Leu Ser Trp Thr Asp Asp Glu Thr Asp Pro Gln Tyr Glu Arg Ser Val Glu Ala Leu Ser Val Leu Ser Asn Ser Ile Asp Ala Arg Gly Arg Lys Ile Gln Val Ile Lys Leu Tyr Ile Pro Glu Pro Leu Tyr Met Thr Glu Glu Glu Ser Ser Gly Ile Thr Gln Asp Gly Glu Ala Ile Pro Arq Leu Ala Gly Thr Arg Leu Ala Ala Ser Tyr Val Asn Phe Tyr Ile Ala Asn Gly Gly Ile Ile Ala Pro Gln Phe Gly Asp Pro Ile Arg Asp Lys Glu Ala Ile Arg Val Leu Ser Asp Thr Phe Pro His His Ser Val Val Gly Ile Glu Asn Ala Arg Glu Ile Val Leu Ala Gly Gly Asn Ile His Cys Ile Thr Gln Gln Pro Ala Glu Pro Thr Ser Val Ala Glu Asn Gly His

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155 160

480

tcc atc aat ttt cta gaa tct tcc act tca tat gct gct cct aca tgg

Ser Ile Asn Phe Leu Glu Ser Ser Thr Ser Tyr Ala Ala Pro Thr Trp

150

145

gga Gly	ttt Phe	gga Gly	ccc Pro	aat Asn 165	tct Ser	gac Asp	aag Lys	ctg Leu	agg Arg 170	tat Tyr	ggt Gly	tca Ser	ctg Leu	cca Pro 175	cgt Arg	528
gaa Glu	gct Ala	gtt Val	tgt Cys 180	act Thr	gag Glu	aac Asn	ttg Leu	acc Thr 185	cca Pro	tgg Trp	cta Leu	aag Lys	tta Leu 190	ctt Leu	cct Pro	576
tgt Cys	aga Arg	gat Asp 195	aag Lys	gat Asp	ggt Gly	att Ile	tct Ser 200	gcg Ala	tta Leu	atg Met	aat Asn	agg Arg 205	cca Pro	tct Ser	gtt Val	624
tac Tyr	aga Arg 210	Gly ggg	ttt Phe	tat Tyr	cat His	tct Ser 215	cag Gln	aga Arg	ttg Leu	cat His	tta Leu 220	tcc Ser	acg Thr	gtt Val	gaa Glu	672
tct Ser 225	ggt Gly	caa Gln	gag Glu	gga Gly	ttg Leu 230	ggt Gly	tct Ser	ggt Gly	ata Ile	gtg Val 235	ctg Leu	gag Glu	cag Gln	acg Thr	ctt Leu 240	720
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ctt Leu	ggt Gly 290	tac Tyr	gaa Glu	tca Ser	aaa Lys	aac Asn 295	gtg Val	gat Asp	aca Thr	gaa Glu	ata Ile 300	gaa Glu	gca Ala	cac His	caa Gln	912
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Gly	Ser 370	Gly	aac Asn	Glu	Arg	Gly 375	Ala	Ile	Ala	Ile	Leu 380	Leu	Lys	Ala	Thr	1152
gaa Glu 385	Ser	cag Gln	gag Glu	aag Lys	tta Leu 390	Ser	ggc	aga Arg	gat Asp	ctc Leu 395	Thr	aat Asn	ggc Gly	caa Gln	tgt Cys 400	1200

			gca		_4_	+ +a	~~ ~	2++	++0	cca	taa	tat	att	aad	att	-	1248
aca Thr	Ile	aaa Lys	Ala	Asn 405	Ile	Phe	Gln	Ile	Phe 410	Pro	Trp	Tyr	Ile	Lys 415	Val		
tat Tyr	tat Tyr	cat His	act Thr 420	cta Leu	caa Gln	atc Ile	ttt Phe	gtg Val 425	gat Asp	caa Gln	caa Gln	cag Gln	aag Lys 430	aca Thr	gac Asp	` ;	1296
agt Ser	gag Glu	gtc Val 435	tta Leu	aag Lys	aag Lys	atc Ile	aat Asn 440	gtc Val	tca Ser	cca Pro	tct Ser	acg Thr 445	gat Asp	aag Lys	gtg Val	;	1344
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agc Ser	aac Asn	tcg Ser 515	ccc Pro	tta Leu	tta Leu	tca Ser	agt Ser 520	tta Leu	aag Lys	gaa Glu	aaa Lys	tcc Ser 525	tta Leu	gta Val	cgc Arg		1584
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atg Met 545	cct Pro	tac Tyr	aac Asn	gta Val	atc Ile 550	acg Thr	atc Ile	aca Thr	tgc Cys	acc Thr 555	atc Ile	ttc Phe	gca Ala	ttg Leu	tat Tyr 560		1680
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agg Arg	ttt Phe	ctc Leu	aaa Lys 580	agc Ser	caa Gln	gca Ala	gga Gly	aag Lys 585	aaa Lys	aca Thr	ggt Gly	gj aaa	ctt Leu 590	Lys	cag Gln		1776
tta Leu	tta Leu	tcg Ser 595	Arg	atc Ile	aca Thr	gcc Ala	aag Lys 600	Ile	aga Arg	gly aaa	aga Arg	cca Pro 605	Ile	gaa Glu	gca Ala		1824
cca Pro	tca Ser 610	Ser	tca Ser	gaa Glu	gct Ala	gaa Glu 615	Ser	tcg Ser	gtc Val	ttg Leu	Ser 620	Ser	aaa Lys	ctt Leu	atc Ile		1872
tta Leu 625	Lys	atc Ile	ata Ile	tta Leu	gtt Val 630	Ala	gga Gly	gct Ala	gct Ala	gca Ala 635	Ala	tgg Trp	caa Gln	tat Tyr	ttt Phe 640		1920

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Phe His Phe Glu Asn Arg Ala Pro Pro Ser Asn Ser His Gly Arg His 50 55 60

His His Leu Phe Pro Lys Ala Ile Ser Gln Leu Val Gln Lys Phe Arg 65 70 75 80

Val Lys Glu Met Glu Leu Ser Phe Thr Gln Gly Arg Trp Asn His Glu 85 90 95

His Trp Gly Gly Phe Asp Pro Leu Ser Ser Met Asn Ala Lys Pro Val 100 105 110

Gly Val Glu Leu Trp Ala Val Phe Asp Val Pro Gln Ser Gln Val Asp 115 120 125

Thr Ser Trp Lys Asn Leu Thr His Ala Leu Ser Gly Leu Phe Cys Ala 130 135 140

Ser Ile Asn Phe Leu Glu Ser Ser Thr Ser Tyr Ala Ala Pro Thr Trp 145 150 155 160

Gly Phe Gly Pro Asn Ser Asp Lys Leu Arg Tyr Gly Ser Leu Pro Arg 165 170 175

Glu Ala Val Cys Thr Glu Asn Leu Thr Pro Trp Leu Lys Leu Leu Pro 180 185 190

- Cys Arg Asp Lys Asp Gly Ile Ser Ala Leu Met Asn Arg Pro Ser Val 195 200 205
- Tyr Arg Gly Phe Tyr His Ser Gln Arg Leu His Leu Ser Thr Val Glu 210 215 220
- Ser Gly Gln Glu Gly Leu Gly Ser Gly Ile Val Leu Glu Gln Thr Leu 225 230 235 240
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- Pro Ser Trp Ser Leu Ser Ser Leu Phe Gly Arg Gln Val Val Gly Arg 260 265 270
- Cys Val Leu Ala Lys Ser Ser Asn Val Tyr Leu Gln Leu Glu Gly Leu 275 280 285
- Leu Gly Tyr Glu Ser Lys Asn Val Asp Thr Glu Ile Glu Ala His Gln 290 295 300
- Leu Trp Lys Asn Ala Glu Phe Glu Leu Ser Leu Lys Pro Glu Arg Val 305 310 315
- Ile Arg Glu Ser Cys Ser Phe Leu Phe Ile Phe Asp Ile Asp Lys Ser 325 330 335
- Ser Asp Ser Glu Pro Phe Asp Leu Gly Leu Thr Trp Lys Arg Pro Ser 340 345 350
- Lys Trp Ser Cys Gln Gln Ala Pro Leu His Ser Ser Arg Phe Leu Met 355 360 365
- Gly Ser Gly Asn Glu Arg Gly Ala Ile Ala Ile Leu Leu Lys Ala Thr 370 375 380
- Glu Ser Gln Glu Lys Leu Ser Gly Arg Asp Leu Thr Asn Gly Gln Cys 385 390 395
- Thr Ile Lys Ala Asn Ile Phe Gln Ile Phe Pro Trp Tyr Ile Lys Val

Tyr Tyr His Thr Leu Gln Ile Phe Val Asp Gln Gln Gln Lys Thr Asp 420 425 430

Ser Glu Val Leu Lys Lys Ile Asn Val Ser Pro Ser Thr Asp Lys Val 435 440 445

Ser Ser Gly Met Met Glu Met Met Leu Glu Leu Pro Cys Glu Val Lys 450 455 460

Ser Val Ala Ile Ser Ile Glu Tyr Asp Lys Gly Phe Leu His Ile Asp 465 470 475 480

Glu Tyr Pro Pro Asp Ala Asn Gln Gly Phe Asp Ile Pro Ser Ala Leu 485 490 495

Ile Ser Phe Pro Asp His His Ala Ser Leu Asp Phe Gln Glu Glu Leu 500 505 510

Ser Asn Ser Pro Leu Leu Ser Ser Leu Lys Glu Lys Ser Leu Val Arg 515 520 525

Ser Tyr Thr Glu Val Leu Leu Val Pro Leu Thr Thr Pro Asp Phe Ser 530 535 540

Met Pro Tyr Asn Val Ile Thr Ile Thr Cys Thr Ile Phe Ala Leu Tyr 545 550 555 560

Phe Gly Ser Leu Leu Asn Val Leu Arg Arg Arg Ile Gly Glu Glu 565 570 575

Arg Phe Leu Lys Ser Gln Ala Gly Lys Lys Thr Gly Gly Leu Lys Gln 580 585 590

Leu Leu Ser Arg Ile Thr Ala Lys Ile Arg Gly Arg Pro Ile Glu Ala 595 600 605

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90

85

95

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aaa Lys	tta Leu 130	gta Val	gaa Glu	tct Ser	cct Pro	tac Tyr 135	gtt Val	gat Asp	tct Ser	ata Ile	gat Asp 140	tgg Trp	gca Ala	agg Arg	tgg Trp	432
cat His 145	ttt Phe	ttc Phe	tgg Trp	gtt Val	gac Asp 150	gag Glu	aga Arg	gtt Val	gtt Val	ccc Pro 155	aag Lys	aat Asn	cac His	gat Asp	gat Asp 160	480
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Arg Lys Val Lys Ser Met Ala Thr Thr Asn Ile Gly Lys Glu Glu Lys 65 70 75 80

Lys Arg Val Glu Ile Tyr Asp Leu Glu Glu Asn Leu Val Ile Asp Leu 85 90 95

Ala Lys Phe Thr Ala Asp Leu Ser Asp Lys Phe Cys Lys Glu Arg Gly 100 105 110

Ala Phe Thr Val Val Val Ser Gly Gly Ser Leu Ile Lys Ser Leu Arg 115 120 125

Lys Leu Val Glu Ser Pro Tyr Val Asp Ser Ile Asp Trp Ala Arg Trp 130 135 140

His Phe Phe Trp Val Asp Glu Arg Val Val Pro Lys Asn His Asp Asp 145 150 155 160

Ser Asn Tyr Lys Leu Ala Tyr Asp Ser Phe Leu Ser Lys Val Pro Ile 165 170 175

Pro Pro Gly Asn Val Tyr Ala Ile Asn Glu Ala Leu Ser Ala Glu Ala 180 185 190

Ala Ala Asp Asp Tyr Glu Thr Cys Leu Lys His Leu Val Asn Thr Asn 195 200 205

Ile Leu Arg Val Ser Glu Ser Thr Gly Phe Pro Lys Phe Asp Leu Met 210 215 220

Leu Leu Gly Met Gly Pro Asp Gly His Val Ala Ser Leu Phe Pro Gly 225 230 235

His Gly Leu Cys Asn Glu Ser Lys Lys Trp Val Val Ser Ile Ser Asp 245 250 255

Ser Pro Lys Pro Pro Ser Glu Arg Ile Thr Phe Thr Phe Pro Val Ile 260 265 270

Asn Ser Ser Ala His Val Ala Leu Val Val Cys Gly Ser Gly Lys Ala 275 280 285

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ţ	tt Phe	ctc Leu	cca Pro 35	cac His	ggc Gly	ggc Gly	gct Ala	tta Leu 40	aga Arg	acc Thr	ggc Gly	gtt Val	tcg Ser 45	tgt Cys	agc Ser	tgg Trp	144
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gtg Val	aat Asn	gaa Glu 755	gga Gly	gat Asp	gac Asp	aaa Lys	agc Ser 760	gga Gly	gaa Glu	aca Thr	gag Glu	gta Val 765	gtt Val	gaa Glu	cca Pro	23
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Val 65	Ala	Glu	Lys	Glu	Thr 70	Thr	Glu	Glu	Gly	Ser 75	Gly	Glu	Lys	Phe	Glu 80	
Tyr	Gln	Ala	Glu	Val 85	Ser	Arg	Leu	Leu	Asp 90	Leu	ılle	Val	His	Ser 95	Leu	
Tyr	Ser	His	Lys		. Val	Phe	Leu	Arg 105		Leu	val	Ser	Asn	Ala	Ser	

Asp Ala Leu Asp Lys Leu Arg Phe Leu Ser Val Thr Glu Pro Ser Leu

Leu Gly Asp Gly Gly Asp Leu Glu Ile Arg Ile Lys Pro Asp Pro Asp Asn Gly Thr Ile Thr Ile Thr Asp Thr Gly Ile Gly Met Thr Lys Glu Glu Leu Ile Asp Cys Leu Gly Thr Ile Ala Gln Ser Gly Thr Ser Lys Phe Leu Lys Ala Leu Lys Glu Asn Lys Asp Leu Gly Ala Asp Asn Gly Leu Ile Gly Gln Phe Gly Val Gly Phe Tyr Ser Ala Phe Leu Val Ala Glu Lys Val Val Val Ser Thr Lys Ser Pro Lys Ser Asp Lys Gln Tyr Val Trp Glu Ser Val Ala Asp Ser Ser Ser Tyr Leu Ile Arg Glu Glu Thr Asp Pro Asp Asn Ile Leu Arg Arg Gly Thr Gln Ile Thr Leu Tyr Leu Arg Glu Asp Asp Lys Tyr Glu Phe Ala Glu Ser Thr Arg Ile Lys Asn Leu Val Lys Asn Tyr Ser Gln Phe Val Gly Phe Pro Ile Tyr Thr Trp Gln Glu Lys Ser Arg Thr Ile Glu Val Glu Glu Asp Glu Pro Val Lys Glu Gly Glu Gly Glu Pro Lys Lys Lys Thr Thr Lys Thr Glu Lys Tyr Trp Asp Trp Glu Leu Ala Asn Glu Thr Lys Pro Leu Trp Met Arg Asn Ser Lys Glu Val Glu Lys Gly Glu Tyr Asn Glu Phe Tyr Lys Lys Ala Phe Asn Glu Phe Leu Asp Pro Leu Ala His Thr His Phe

Thr Thr Glu Gly Glu Val Glu Phe Arg Ser Ile Leu Tyr Ile Pro Gly 370 375 380

Met Gly Pro Leu Asn Asn Glu Asp Val Thr Asn Pro Lys Thr Lys Asn 385 395 400

Ile Arg Leu Tyr Val Lys Arg Val Phe Ile Ser Asp Asp Phe Asp Gly 405 410 415

Glu Leu Phe Pro Arg Tyr Leu Ser Phe Val Lys Gly Val Val Asp Ser 420 425 430

Asp Asp Leu Pro Leu Asn Val Ser Arg Glu Ile Leu Gln Glu Ser Arg 435 440 445

Ile Val Arg Ile Met Arg Lys Arg Leu Ile Arg Lys Thr Phe Asp Met 450 455 460

Ile Gln Glu Ile Ser Glu Ser Glu Asn Lys Glu Asp Tyr Lys Lys Phe 465 470 475 480

Trp Glu Asn Phe Gly Arg Phe Leu Lys Leu Gly Cys Ile Glu Asp Thr 485 490 495

Gly Asn His Lys Arg Ile Thr Pro Leu Leu Arg Phe Phe Ser Ser Lys 500 505 510

Asn Glu Glu Leu Thr Ser Leu Asp Asp Tyr Ile Glu Asn Met Gly 515 520 525

Glu Asn Gln Lys Ala Ile Tyr Tyr Leu Ala Thr Asp Ser Leu Lys Ser 530 540

Ala Lys Ser Ala Pro Phe Leu Glu Lys Leu Ile Gln Lys Asp Ile Glu 545 550 555 560

Val Leu Tyr Leu Val Glu Pro Ile Asp Glu Val Ala Ile Gln Asn Leu 565 570 575

Gln Thr Tyr Lys Glu Lys Lys Phe Val Asp Ile Ser Lys Glu Asp Leu 580 585 590

Glu Leu Gly Asp Glu Asp Glu Val Lys Asp Arg Glu Ala Lys Gln Glu 595 600 605

Phe Asn Leu Leu Cys Asp Trp Ile Lys Gln Gln Leu Gly Asp Lys Val 610 615 620

Ala Lys Val Gln Val Ser Asn Arg Leu Ser Ser Ser Pro Cys Val Leu 625 630 635 640

Val Ser Gly Lys Phe Gly Trp Ser Ala Asn Met Glu Arg Leu Met Lys 645 650 655

Ala Gln Ala Leu Gly Asp Thr Ser Ser Leu Glu Phe Met Arg Gly Arg 660 665 670

Arg Ile Leu Glu Ile Asn Pro Asp His Pro Ile Ile Lys Asp Leu Asn 675 680 685

Ala Ala Cys Lys Asn Ala Pro Glu Ser Thr Glu Ala Thr Arg Val Val 690 695 700

Asp Leu Leu Tyr Asp Thr Ala Ile Ile Ser Ser Gly Phe Thr Pro Asp 705 710 715 720

Ser Pro Ala Glu Leu Gly Asn Lys Ile Tyr Glu Met Met Ala Met Ala 725 730 735

Val Gly Gly Arg Trp Gly Arg Val Glu Glu Glu Glu Glu Ser Ser Thr 740 745 750

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aag Lys	ctt Leu	ggt Gly	ttt Phe	gtg Val 85	agg Arg	act Thr	ttg Leu	ttg Leu	att Ile 90	gat Asp	aat Asn	tat Tyr	gat Asp	agt Ser 95	tat Tyr	2	:88
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ctt Leu	gaa Glu	tgc Cys	cgt Arg	gat Asp 165	atc Ile	cca Pro	att Ile	cta Leu	ggc Gly 170	gtc Val	tgc Cys	ctt Leu	ggc	cac His 175	cag Gln	5	528
gca Ala	cta Leu	ggt Gly	tat Tyr 180	gtc Val	cat His	gga Gly	gct Ala	cat His 185	gtg Val	gtg Val	cat His	gcc Ala	ccg Pro 190	gaa Glu	cca Pro	5	576
gtc Val	cat His	gga Gly 195	cgg Arg	ttg Leu	agt Ser	gly aaa	att Ile 200	gaa Glu	cat His	gat Asp	GJA aaa	aac Asn 205	ata Ile	ttg Leu	ttt Phe	€	524
tct Ser	gat Asp 210	Ile	cca Pro	tcc Ser	gly aaa	aga Arg 215	aac Asn	tct Ser	gat Asp	ttt Phe	aag Lys 220	gtt Val	gtt Val	aga Arg	tac Tyr	€	572

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ata Ile	gcg Ala	tgg Trp	acg Thr	att Ile 245	tat Tyr	gat Asp	gac Asp	act Thr	ggc Gly 250	tct Ser	ttc Phe	tct Ser	gag Glu	aag Lys 255	aat Asn	768
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gtc Val	att Ile	cct Pro 275	gtt Val	tca Ser	gaa Glu	aag Lys	tta Leu 280	gaa Glu	aat Asn	cga Arg	agt Ser	cat His 285	tgg Trp	cct Pro	tcg Ser	864
tcc Ser	cat His 290	gtt Val	aat Asn	Gly 999	aaa Lys	caa Gln 295	gat Asp	aga Arg	cac His	att Ile	ctc Leu 300	atg Met	ggc	atc Ile	atg Met	912
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Val	Arg	Asn 435		Phe	Met	Glu	Leu 440	Phe	Gly	Lys	Asn	Arg 445	Gly	Asn	Asp	1344
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ctg Leu	att Ile	gaa Glu	gat Asp 500	tct Ser	cag Gln	agt Ser	tct Ser	act Thr 505	gag Glu	aaa Lys	caa Gln	ttc Phe	ttg Leu 510	gaa Glu	gaa Glu	15	36
Gly	ttt Phe	ctt Leu 515	gat Asp	ttt Phe	ctc Leu	cgt Arg	aag Lys 520	gag Glu	ctt Leu	tca Ser	tct Ser	atc Ile 525	tct Ser	tat Tyr	gat Asp	15	84
gag Glu	aag Lys 530	gac Asp	ttc Phe	gaa Glu	gag Glu	ttg Leu 535	cct Pro	ttt Phe	gat Asp	ttt Phe	tgc Cys 540	ggt Gly	gga Gly	tac Tyr	gta Val	16	32
ggt Gly 545	tgt Cys	att Ile	ejå aaa	tat Tyr	gat Asp 550	att Ile	aaa Lys	gtg Val	gaa Glu	tgt Cys 555	gga Gly	atg Met	cca Pro	att Ile	aat Asn 560	16	80
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aga Arg	gag Glu 690	Arg	aat Asn	cca Pro	gca Ala	cca Pro 695	Tyr	gca Ala	gca Ala	ttt Phe	ctc Leu 700	Asn	ttc Phe	tca Ser	aat Asn	21	L12

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cgt Arg	Gly	tcc Ser	acg Thr 740	cct Pro	gaa Glu	gaa Glu	gat Asp	gaa Glu 745	ttt Phe	ctt Leu	aaa Lys	ttg Leu	caa Gln 750	ttg Leu	aaa Lys	2256
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Thr Arg Lys Val Leu Ala Ser Ser Arg Tyr Val Pro Gly Lys Leu Glu 50 55 60

Asp Leu Ser Val Val Lys Lys Ser Leu Pro Arg Arg Glu Pro Val Glu 65 70 75 80

Lys Leu Gly Phe Val Arg Thr Leu Leu Ile Asp Asn Tyr Asp Ser Tyr 85 90 95

Thr Phe Asn Ile Tyr Gln Ala Leu Ser Thr Ile Asn Gly Val Pro Pro 100 105 110

Val Val Ile Arg Asn Asp Glu Trp Thr Trp Glu Glu Ala Tyr His Tyr 115 120 125

Leu Tyr Glu Asp Val Ala Phe Asp Asn Ile Val Ile Ser Pro Gly Pro 130 135 140

Gly Ser Pro Met Cys Pro Ala Asp Ile Gly Ile Cys Leu Arg Leu Leu 145 150 155 160

Leu Glu Cys Arg Asp Ile Pro Ile Leu Gly Val Cys Leu Gly His Gln 165 170 175

Ala Leu Gly Tyr Val His Gly Ala His Val Val His Ala Pro Glu Pro 180 185 190

Val His Gly Arg Leu Ser Gly Ile Glu His Asp Gly Asn Ile Leu Phe 195 200 205

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Thr Phe Trp Leu Asp Thr Ser Ser Ser Asp Lys Ala Arg Gly Arg Phe Ser Phe Met Gly Gly Lys Gly Gly Ser Leu Trp Lys Gln Leu Thr Phe Ser Leu Ser Asp Gln Ser Glu Val Thr Ser Lys His Ala Gly His Leu Leu Ile Glu Asp Ser Gln Ser Ser Thr Glu Lys Gln Phe Leu Glu Glu Gly Phe Leu Asp Phe Leu Arg Lys Glu Leu Ser Ser Ile Ser Tyr Asp Glu Lys Asp Phe Glu Glu Leu Pro Phe Asp Phe Cys Gly Gly Tyr Val Gly Cys Ile Gly Tyr Asp Ile Lys Val Glu Cys Gly Met Pro Ile Asn Arg His Lys Ser Asn Ala Pro Asp Ala Cys Phe Phe Phe Ala Asp Asn Val Val Ala Ile Asp His Gln Leu Asp Asp Val Tyr Ile Leu Ser Leu Tyr Glu Glu Gly Thr Ala Glu Thr Ser Phe Leu Asn Asp Thr Glu Glu Lys Leu Ile Ser Leu Met Gly Leu Ser Thr Arg Lys Leu Glu Asp Gln Thr Leu Pro Val Ile Asp Ser Ser Gln Ser Lys Thr Ser Phe Val Pro Asp Lys Ser Arg Glu Gln Tyr Ile Asn Asp Val Gln Ser Cys Met Lys Tyr Ile Lys Asp Gly Glu Ser Tyr Glu Leu Cys Leu Thr Thr Gln Asn Arg Arg Lys Ile Gly Asn Ala Asp Pro Leu Gly Leu Tyr Leu His Leu

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ctg Leu	aaa Lys	ctg Leu	aga Arg 340	atc Ile	atc Ile	gaa Glu	cat His	aat Asn 345	atc Ile	ctc Leu	gtt Val	gtc Val	tca Ser 350	aag Lys	tac Tyr	1056
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1248

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1296 aaq ctt cta gat ctt gtg gaa aag agt tgc cac caa att cac aag gaa Lys Leu Leu Asp Leu Val Glu Lys Ser Cys His Gln Ile His Lys Glu 425 420

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<212> PRT

<213> Arabidopsis thaliana

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Leu Leu Asn Glu Gln Ile Leu Asn Leu Ser Lys Lys Arg Gly Gln Leu

Lys Gln Ala Val Gln Ser Met Val Gln Gln Ala Met Gln Tyr Ile Asp 65

Gln Thr Pro Asp Ile Glu Thr Arg Ile Glu Leu Ile Lys Thr Leu Asn 90 85

Asn Val Ser Ala Gly Lys Ile Tyr Val Glu Ile Glu Arg Ala Arg Leu 110 105 100

Thr Lys Lys Leu Ala Lys Ile Lys Glu Glu Gln Gly Gln Ile Ala Glu 125 115 120

PCT/EP02/07929 **WO** 03/008440

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Ile Pro Asp Phe Lys Met Leu Leu Lys Gln Val Val Thr Met Glu Val

Ile Gln Trp Thr Ser Leu Trp Asn Lys Tyr Lys Asp Glu Phe Glu Lys

Glu Lys Ser Met Ile Gly Gly Ser Leu Gly Asp Lys Ala Gly Glu Asp

Leu Lys Leu Arg Ile Ile Glu His Asn Ile Leu Val Val Ser Lys Tyr

Tyr Ala Arg Ile Thr Leu Lys Arg Leu Ala Glu Leu Leu Cys Leu Ser

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ttc gct ggg Phe Ala Gly 115	ctt tct Leu Ser	gag att Glu Ile	atg (Met (gag att Glu Ile	cct Pro	gtg Val	ctt Leu 125	aaa Lys	gga Gly	gaa Glu
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120

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- Ser Lys Met Pro Thr Leu Glu Glu Tyr Gly Thr Asn Leu Thr Lys Leu 275 280 285
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Ala Gln Arg Ile Ala Ser Gly Asp Val Pro Glu Thr Ile Glu Gly Lys 340 345 350

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Arg Gly Glu Phe Glu Glu Arg Leu Lys Lys Leu Met Glu Glu Ile Arg 370 375 380

Gln Ser Asp Glu Ile Ile Leu Phe Ile Asp Glu Val His Thr Leu Ile 385 390 395 400

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Pro Ala Leu Ala Arg Gly Glu Leu Gln Cys Ile Gly Ala Thr Thr Ile 420 425 430

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cta cg Leu Ar 21	g Asr	tat Tyr	ttg Leu	cat His	tac Tyr 215	aac Asn	ctc Leu	tat Tyr	gat Asp	cag Gln 220	gca Ala	gag Glu	aag Lys	cta Leu	672
aga to	a aag	gca	cct	cgc	ttt	gag	gct	cat	tca	aac	caa	cag	ttt	tgt	720

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acg Thr	gac Asp	gca Ala	aaa Lys 260	gag Glu	agc Ser	ctt Leu	ctt Leu	cag Gln 265	gcg Ala	gcc Ala	agg Arg	aaa Lys	gcc Ala 270	cct Pro	ata Ile	816
gca Ala	gct Ala	ttg Leu 275	ggc Gly	ttc Phe	agg Arg	atc Ile	caa Gln 280	tgc Cys	aat Asn	aaa Lys	tgg Trp	gca Ala 285	att Ile	ctg Leu	gtt Val	864
cgt Arg	cta Leu 290	ctg Leu	ctg Leu	ggt Gly	gag Glu	ata Ile 295	cca Pro	gag Glu	cgt Arg	tct Ser	atc Ile 300	ttc Phe	act Thr	caa Gln	aag Lys	912
ggt Gly 305	atg Met	gag Glu	aag Lys	gcc Ala	ctc Leu 310	aga Arg	ccc Pro	tac Tyr	ttc Phe	gag Glu 315	cta Leu	aca Thr	aat Asn	gcg Ala	gtt Val 320	960
agg Arg	att Ile	gly aaa	gac Asp	ttg Leu 325	gag Glu	ttg Leu	ttt Phe	agg Arg	aca Thr 330	gtc Val	cag Gln	gag Glu	aag Lys	ttc Phe 335	ttg Leu	1008
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cac His	aat Asn	gtc Val 355	atc Ile	agg Arg	act Thr	gga Gly	ctg Leu 360	cgg Arg	aac Asn	ata Ile	agt Ser	atc Ile 365	tcc Ser	tac Tyr	tca Ser	1104
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aac Asn 385	cct Pro	gtg Val	gct Ala	gat Asp	gcg Ala 390	gaa Glu	agc Ser	atc Ile	gtg Val	gca Ala 395	aag Lys	gcc Ala	ata Ile	cgc Arg	gac Asp 400	1200
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aaa Lys	gaa Glu	act Thr	999 Gly 420	Asp	atc Ile	tac Tyr	tcg Ser	acg Thr 425	Asn	gag Glu	cca Pro	caa Gln	act Thr 430	gcg Ala	ttc Phe	1296
aac Asn	tca Ser	aga Arg 435	att Ile	gct Ala	ttc Phe	tgc Cys	ctc Leu 440	Asn	atg Met	cat His	aac Asn	gaa Glu 445	gct Ala	gtc Val	aga Arg	1344
gca Ala	ttg Leu 450	Arg	ttt Phe	cct	cct Pro	aac Asn 455	Thr	cac His	aag Lys	gag Glu	aaa Lys 460	Glu	agc Ser	gat Asp	gag Glu	1392
aag	agg	aga	gag	agg	aag	caa	cag	gaa	gaa	gag	ctt	gct	aag	cat	atg	1440

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<213> Arabidopsis thaliana

<400> 84

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Arg Ala Val Arg Leu Thr Ile Gly Leu Arg Gln Lys Leu Thr Gly Ser 50 55 60

Val Leu Ser Ser Phe Leu Asp Phe Ala Leu Val Pro Gly Ser Glu Ala 65 70 75 80

His Ser Arg Leu Ser Ser Phe Val Pro Lys Gly Asp Glu His Asp Met 85 90 95

Glu Val Asp Thr Ala Ser Ser Ala Thr Gln Ala Ala Pro Ser Lys His 100 105 110

Leu Pro Ala Glu Leu Glu Ile Tyr Cys Tyr Phe Ile Val Leu Leu Phe 115 120 125

Leu Ile Asp Gln Lys Lys Tyr Asn Glu Ala Lys Ala Cys Ser Ser Ala 130 135 140

Ser Ile Ala Arg Leu Lys Asn Val Asn Arg Arg Thr Ile Asp Val Ile 145 150 155 160

Ala Ser Arg Leu Tyr Phe Tyr Tyr Ser Leu Ser Tyr Glu Gln Thr Gly
165 170 175

Asp Leu Ala Glu Ile Arg Gly Thr Leu Leu Ala Leu His His Ser Ala 180 185 190

Thr Leu Arg His Asp Glu Leu Gly Gln Glu Thr Leu Leu Asn Leu Leu 195 200 205

Leu Arg Asn Tyr Leu His Tyr Asn Leu Tyr Asp Gln Ala Glu Lys Leu 210 215 220

Arg Ser Lys Ala Pro Arg Phe Glu Ala His Ser Asn Gln Gln Phe Cys 225 230 235 240

Arg Tyr Leu Phe Tyr Leu Gly Lys Ile Arg Thr Ile Gln Leu Glu Tyr 245 250 255

Thr Asp Ala Lys Glu Ser Leu Leu Gln Ala Ala Arg Lys Ala Pro Ile 260 265 270

Ala Ala Leu Gly Phe Arg Ile Gln Cys Asn Lys Trp Ala Ile Leu Val 275 280 285

Arg Leu Leu Gly Glu Ile Pro Glu Arg Ser Ile Phe Thr Gln Lys 290 295 300

Gly Met Glu Lys Ala Leu Arg Pro Tyr Phe Glu Leu Thr Asn Ala Val 305 310 315 320

Arg Ile Gly Asp Leu Glu Leu Phe Arg Thr Val Gln Glu Lys Phe Leu 325 330 335

Asp Thr Phe Ala Gln Asp Arg Thr His Asn Leu Ile Val Arg Leu Arg 340 345 350

His Asn Val Ile Arg Thr Gly Leu Arg Asn Ile Ser Ile Ser Tyr Ser 355 360 365

Arg Ile Ser Leu Pro Asp Val Ala Lys Lys Leu Arg Leu Asn Ser Glu 370 380

Asn Pro Val Ala Asp Ala Glu Ser Ile Val Ala Lys Ala Ile Arg Asp 385 390 395 400

Gly	Ala	Ile	Asp	Ala 405	Thr	Ile	Asp	His	Lys 410	Asn	Gly	Cys	Met	Val 415	Ser	
Lys	Glu	Thr	Gly 420	Asp	Ile	Tyr	Ser	Thr 425	Asn	Glu	Pro	Gln	Thr 430	Ala	Phe	
Asn	Ser	Arg 435	Ile	Ala	Phe	Cys	Leu 440	Asn	Met	His	Asn	Glu 445	Ala	Val	Arg	
Ala	Leu 450	Arg	Phe	Pro		Asn 455	Thr	His	Lys	Glu	Lуs 460	Glu	Ser	Asp	Glu	
Lys 465	Arg	Arg	Glu	Arg	Lys 470	Gln	Gln	Glu	Glu	Glu 475	Leu	Ala	Lys	His	Met 480	
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aaa Lys	gcc Ala	aaa Lys	atg Met 20	gcc Ala	tcc Ser	atg Met	ato Ile	gat Asp 25	cag Gln	ctt Leu	cag Gln	cto Leu	cgt Arg 30	gat Asp	agt Ser	96
ttg Lev	agg Arg	atg Met 35	tac Tyr	aat Asn	tca Ser	ttg Leu	gtg Val 40	gag Glu	agg Arg	tgt Cys	tto Phe	gtg Val 45	gac Asp	tgt Cys	gtt Val	144
gat Asp													_			192

atg cgt tgc gct gag aag ttc ctt aag cat acg atg cgt gtt ggt atg

Met Arg Cys Ala Glu Lys Phe Leu Lys His Thr Met Arg Val Gly Met
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<211> 93

<212> PRT

<213> Arabidopsis thaliana

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Asp Ser Phe Thr Arg Lys Ser Leu Gln Lys Gln Glu Glu Thr Cys Val 50 55 60

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Arg Phe Ala Glu Leu Asn Gln Asn Ala Pro Thr Gln Asp 85 90

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<223> 70913

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cat His	gct Ala	gtt Val	cag Gln 20	cag Gln	ccg Pro	atg Met	atg Met	tat Tyr 25	gca Ala	gag Glu	ccc Pro	tgg Trp	tgg Trp 30	aaa Lys	aac Asn		96
aac Asn	tcc Ser	ttt Phe 35	ggt Gly	gtt Val	gta Val	cct Pro	caa Gln 40	gcg Ala	aga Arg	cct Pro	tct Ser	gga Gly 45	att Ile	cca Pro	tca Ser	•	144
aat Asn	tcc Ser 50	tct Ser	tct Ser	ttg Leu	gat Asp	tgc Cys 55	ccc Pro	aat Asn	ggt Gly	tcc Ser	gag Glu 60	tca Ser	aac Asn	gat Asp	gtt Val		192
cat His 65	tca Ser	gca Ala	tct Ser	gaa Glu	gac Asp 70	ggt Gly	gcg Ala	ttg Leu	aat Asn	ggt Gly 75	gaa Glu	aac Asn	gat Asp	ggc Gly	act Thr 80		240
tgg Trp	aag Lys	gat Asp	tca Ser	caa Gln 85	gct Ala	gca Ala	act Thr	tcc Ser	tct Ser 90	cgt Arg	tca Ser	gat Asp	aat Asn	cac His 95	gga Gly		288
atg Met	gaa Glu	gga Gly	aat Asn 100	gac Asp	cca Pro	gcg Ala	ctc Leu	tct Ser 105	atc Ile	cgt Arg	aac Asn	atg Met	cat His 110	gat Asp	cag Gln		336
cca Pro	ctt Leu	gta Val 115	caa Gln	cca Pro	cca Pro	gag Glu	ctt Leu 120	gtt Val	gga Gly	cac His	tat Tyr	atc Ile 125	gct Ala	tgt Cys	gtc Val		384
cca Pro	aac Asn 130	cca Pro	tat Tyr	cag Gln	gat Asp	cca Pro 135	tat Tyr	tat Tyr	Gly 999	gga Gly	ttg Leu 140	atg Met	gga Gly	gca Ala	tat Tyr		432
ggt Gly 145	cat His	cag Gln	caa Gln	ttg Leu	ggt Gly 150	ttt Phe	cgt Arg	cca Pro	tat Tyr	ctt Leu 155	gga Gly	atg Met	cct Pro	cgt Arg	gaa Glu 160		480
aga Arg	aca Thr	gct Ala	ctg Leu	cca Pro 165	ctt Leu	gac Asp	atg Met	gca Ala	caa Gln 170	gag Glu	ccc Pro	gtt Val	tat Tyr	gtg Val 175	aat Asn		528
gca Ala	aag Lys	cag Gln	tac Tyr 180	gag Glu	gga Gly	att Ile	cta Leu	agg Arg 185	cga Arg	aga Arg	aaa Lys	gca Ala	cgt Arg 190	gcc Ala	aag Lys		576
gca Ala	gag Glu	cta Leu 195	gag Glu	agg Arg	aaa Lys	gtc Val	atc Ile 200	Arg	gac Asp	aga Arg	aag Lys	cca Pro 205	Tyr	ctt Leu	cac His		624
gag	tca	. aga	cac	aag	cat	gca	atg	aga	agg	gca	cga	gcg	agt	gga	ggc		672

Glu Ser Arg His Lys His Ala Met Arg Arg Ala Arg Ala Ser Gly Gly 210 215 220

cgg ttt gcg aag aaa agt gag gta gaa gcg gga gag gat gca gga ggg 720
Arg Phe Ala Lys Lys Ser Glu Val Glu Ala Gly Glu Asp Ala Gly Gly
225 230 235 240

aga gac aga gaa agg ggt tca gca acc aac tca tca ggc tct gaa caa 768
Arg Asp Arg Glu Arg Gly Ser Ala Thr Asn Ser Ser Gly Ser Glu Gln
245 250 255

gtt gag aca gac tct aat gag acc ctg aat tct tct ggt gca cca taa 816
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<210> 88

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His Ala Val Gln Gln Pro Met Met Tyr Ala Glu Pro Trp Trp Lys Asn 20 25 30

Asn Ser Phe Gly Val Val Pro Gln Ala Arg Pro Ser Gly Ile Pro Ser 35. 40 45 -

Asn Ser Ser Ser Leu Asp Cys Pro Asn Gly Ser Glu Ser Asn Asp Val 50 55 60

His Ser Ala Ser Glu Asp Gly Ala Leu Asn Gly Glu Asn Asp Gly Thr 65 70 75 80

Trp Lys Asp Ser Gln Ala Ala Thr Ser Ser Arg Ser Asp Asn His Gly 85 90 95

Met Glu Gly Asn Asp Pro Ala Leu Ser Ile Arg Asn Met His Asp Gln
100 105 110

Pro Leu Val Gln Pro Pro Glu Leu Val Gly His Tyr Ile Ala Cys Val 115 120 125

Pro Asn Pro Tyr Gln Asp Pro Tyr Tyr Gly Gly Leu Met Gly Ala Tyr 130 135 140

Gly His Gln Gln Leu Gly Phe Arg Pro Tyr Leu Gly Met Pro Arg Glu 145 150 155 160

Arg Thr Ala Leu Pro Leu Asp Met Ala Gln Glu Pro Val Tyr Val Asn 165 170 175

Ala Lys Gln Tyr Glu Gly Ile Leu Arg Arg Arg Lys Ala Arg Ala Lys 180 185 190

Ala Glu Leu Glu Arg Lys Val Ile Arg Asp Arg Lys Pro Tyr Leu His 195 200 205

Glu Ser Arg His Lys His Ala Met Arg Arg Ala Arg Ala Ser Gly Gly 210 215 220

Arg Phe Ala Lys Lys Ser Glu Val Glu Ala Gly Glu Asp Ala Gly Gly 225 230 235 240

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<222> (1)..(990)

<223> 71067

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48

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cat His	agg Arg	ctt Leu 35	cgt Arg	cac His	ttt Phe	gag Glu	tgt Cys 40	gaa Glu	ggc	agt Ser	tat Tyr	ccc Pro 45	aag Lys	tat Tyr	cct Pro	144
tat Tyr	ggt Gly 50	tct Ser	ttg Leu	gtc Val	aag Lys	ttt Phe 55	tat Tyr	gca Ala	atg Met	gtg Val	gga Gly 60	ctt Leu	cat His	cgt Arg	tac Tyr	192
aat Asn 65	gtg Val	ttg Leu	gag Glu	gly aaa	aaa Lys 70	aat Asn	ttg Leu	cag Gln	ctc Leu	gat Asp 75	acc Thr	cta Leu	aag Lys	agt Ser	ttc Phe 80	240
aac Asn	atg Met	aga Arg	atc Ile	aat Asn 85	tgt Cys	ggt Gly	gct Ala	tct Ser	tct Ser 90	tac Tyr	tac Tyr	att Ile	act Thr	ttg Leu 95	gct Ala	288
gca Ala	cgc Arg	gtt Val	cca Pro 100	gat Asp	agc Ser	ggt Gly	ttg Leu	aag Lys 105	cag Gln	atc Ile	ttt Phe	cag Gln	gtt Val 110	cta Leu	gtt Val	336
cat His	gaa Glu	gag Glu 115	cgt Arg	ctt Leu	ggc Gly	agt Ser	tta Leu 120	gac Asp	atg Met	aca Thr	tgt Cys	act Thr 125	atc Ile	gct Ala	aga Arg	384
cct Pro	cga Arg 130	gtg Val	act Thr	acc Thr	aat Asn	gtg Val 135	cct Pro	ttt Phe	cta Leu	cgt Arg	ccg Pro 140	cac His	agc Ser	gaa Glu	tca Ser	432
gag Glu 145	tat Tyr	gat Asp	tat Tyr	atg Met	gac Asp 150	aat Asn	gat Asp	gaa Glu	ttg Leu	cct Pro 155	gac Asp	tgg Trp	cct Pro	tca Ser	gag Glu 160	480
att Ile	gct Ala	ttc Phe	gat Asp	gat Asp 165	aca Thr	aaa Lys	cgg Arg	ttt Phe	cat His 170	ctg Leu	gtg Val	aag Lys	gaa Glu	tca Ser 175	gag Glu	528
ttg Leu	cga Arg	gac Asp	aat Asn 180	gat Asp	tgg Trp	att Ile	cga Arg	ctc Leu 185	tat Tyr	ttg Leu	gaa Glu	ctt Leu	aca Thr 190	ctt Leu	gtt Val	576
gct Ala	cac His	gat Asp 195	agg Arg	ttt Phe	ctt Leu	aca Thr	gtt Val 200	cac His	tat Tyr	ctc Leu	tcc Ser	cag Gln 205	Leu	gag Glu	att Ile	624
gtg Val	aaa Lys 210	Val	gcg Ala	att Ile	gaa Glu	gaa Glu 215	gtg Val	gag Glu	caa Gln	ccg Pro	aat Asn 220	Ala	agt Ser	ctc Leu	aac Asn	672
Thr 225	Ьys	Thr	aca Thr	Phe	Val 230	Tyr	Ile	Thr	Tyr	Lys 235	Asp	Leu	Ala	Lys	Ala 240	720
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aat Asn	gag Glu	act Thr	acg Thr 260	gga Gly	ctc Leu	ttg Leu	aga Arg	ctc Leu 265	cgg Arg	ggt Gly	gat Asp	tat Tyr	tgg Trp 270	agt Ser	gga Gly	816
gaa Glu	aga Arg	agt Ser 275	gtg Val	atc Ile	act Thr	ccg Pro	gag Glu 280	gag Glu	gaa Glu	tat Tyr	atg Met	ctt Leu 285	ctc Leu	cat His	ggc Gly	864
gga Gly	gaa Glu 290	aaa Lys	gtt Val	cga Arg	aac Asn	aat Asn 295	gag Glu	cag Gln	cgt Arg	tct Ser	aaa Lys 300	aaa Lys	ctt Leu	aag Lys	cgt Arg	912
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<213> Arabidopsis thaliana

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Tyr Gly Ser Leu Val Lys Phe Tyr Ala Met Val Gly Leu His Arg Tyr 50 55 60

Asn Val Leu Glu Gly Lys Asn Leu Gln Leu Asp Thr Leu Lys Ser Phe 65 70 75 80

Asn Met Arg Ile Asn Cys Gly Ala Ser Ser Tyr Tyr Ile Thr Leu Ala 85 90 95

Ala Arg Val Pro Asp Ser Gly Leu Lys Gln Ile Phe Gln Val Leu Val 100 105 110

His Glu Glu Arg Leu Gly Ser Leu Asp Met Thr Cys Thr Ile Ala Arg 115 120 125

Pro Arg Val Thr Thr Asn Val Pro Phe Leu Arg Pro His Ser Glu Ser 130 135 140

Glu Tyr Asp Tyr Met Asp Asn Asp Glu Leu Pro Asp Trp Pro Ser Glu 145 150 155 160

Ile Ala Phe Asp Asp Thr Lys Arg Phe His Leu Val Lys Glu Ser Glu 165 170 175

Leu Arg Asp Asn Asp Trp Ile Arg Leu Tyr Leu Glu Leu Thr Leu Val 180 185 190

Ala His Asp Arg Phe Leu Thr Val His Tyr Leu Ser Gln Leu Glu Ile 195 200 205

Val Lys Val Ala Ile Glu Glu Val Glu Gln Pro Asn Ala Ser Leu Asn 210 215 220

Thr Lys Thr Thr Phe Val Tyr Ile Thr Tyr Lys Asp Leu Ala Lys Ala 225 230 235 240

Gln Ile Gly Glu Pro Val Asp Arg Lys Ala Ile Val Arg Lys Ile Ile 245 250 255

Asn Glu Thr Thr Gly Leu Leu Arg Leu Arg Gly Asp Tyr Trp Ser Gly 260 265 270

Glu Arg Ser Val Ile Thr Pro Glu Glu Glu Tyr Met Leu Leu His Gly 275 280 285

Gly Glu Lys Val Arg Asn Asn Glu Gln Arg Ser Lys Lys Leu Lys Arg 290 295 300

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<223> 71654

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tct Ser	cag Gln	agg Arg	gat Asp 100	ctt Leu	gcg Ala	tac Tyr	cgt Arg	cgt Arg 105	cgt Arg	act Thr	cgt Arg	act Thr	ggt Gly 110	ttt Phe	gca Ala	33	6
aat Asn	cta Leu	tac Tyr 115	gta Val	aag Lys	aat Asn	ctg Leu	gat Asp 120	agc Ser	tcg Ser	att Ile	act Thr	agc Ser 125	agt Ser	tgc Cys	tta Leu	38	4
gag Glu	cga Arg 130	atg Met	ttt Phe	tgc Cys	ccc Pro	ttt Phe 135	ggt Gly	tcc Ser	ata Ile	ctt Leu	tct Ser 140	tgc Cys	aaa Lys	gtc Val	gtt Val	43	2
gaa Glu 145	gag Glu	aat Asn	ggc Gly	caa Gln	agt Ser 150	aaa Lys	ggt Gly	ttt Phe	ggc Gly	ttt Phe 155	gtt Val	cag Gln	ttt Phe	gat Asp	aca Thr 160	48	0
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Glu	Gln	Ser	Ala	Val	Ser	Ala	Arq	Ser	Ala	Leu	His	Gly	Ser	Met	Val	
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Ser Val His Leu Cys Arg Asn Ser Val Thr Gly Lys Ser Met Cys Tyr 50 55 60

Ala Tyr Ile Asn Phe Asp Ser Pro Phe Ser Ala Ser Asn Ala Met Thr 65 70 75 80

Arg Leu Asn His Ser Asp Leu Lys Gly Lys Ala Met Arg Ile Met Trp 85 90 95

Ser Gln Arg Asp Leu Ala Tyr Arg Arg Arg Thr Arg Thr Gly Phe Ala 100 105 110

Asn Leu Tyr Val Lys Asn Leu Asp Ser Ser Ile Thr Ser Ser Cys Leu 115 120 125

Glu Arg Met Phe Cys Pro Phe Gly Ser Ile Leu Ser Cys Lys Val Val 130 135 140

Glu Glu Asn Gly Gln Ser Lys Gly Phe Gly Phe Val Gln Phe Asp Thr 145 150 155 160

Glu Gln Ser Ala Val Ser Ala Arg Ser Ala Leu His Gly Ser Met Val 165 170 175

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Tyr Gly Thr Val Ser Ser Val Val Val Met Arg Asp Gly Met Gly Arg 225 230 235 240

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Lys Ala Met Glu Ser Leu Cys Gly Leu Gln Leu Gly Ser Lys Lys Leu 260 265 270

Phe Val Gly Lys Ala Leu Lys Lys Asp Glu Arg Arg Glu Met Leu Lys 275 280 285

Gln Lys Phe Ser Asp Asn Phe Ile Ala Lys Pro Asn Met Arg Trp Ser 295 290 Asn Leu Tyr Val Lys Asn Leu Ser Glu Ser Met Asn Glu Thr Arg Leu 310 315 Arg Glu Ile Phe Gly Cys Tyr Gly Gln Ile Val Ser Ala Lys Val Met 330 325 Cys His Glu Asn Gly Arg Ser Lys Gly Phe Gly Phe Val Cys Phe Ser 345 340 Asn Cys Glu Glu Ser Lys Gln Ala Lys Arg Tyr Leu Asn Gly Phe Leu Val Asp Gly Lys Pro Ile Val Val Arg Val Ala Glu Arg Lys Glu Asp 375 Arg Ile Lys Arg Leu Gln Gln Tyr Phe Gln Ala Gln Pro Arg Gln Tyr . 390 385 Thr Gln Ala Pro Ser Ala Pro Ser Pro Ala Gln Pro Val Leu Ser Tyr 410 405 Val Ser Ser Ser Tyr Gly Cys Phe Gln Pro Phe Gln Val Gly Thr Ser 430 420 425 Tyr Tyr Tyr Met Gly Asn Gln Val Pro Gln Met Ser Gly His Gln Asn 440 435 Ile Thr Thr Tyr Val Pro Ala Gly Lys Val Pro Leu Lys Glu Arg Arg Ser Met His Leu Val Tyr Lys His Pro Ala Tyr Pro Val Ala Lys Arg 470 475 465 Gly Ala Lys Gln Thr Leu Val Phe Lys Gly Glu Val Asn Arg Asn Leu 490 485 Glu Ala Ala Thr Cys Ser Lys Ala Thr Thr Ser Glu Glu Asn Arg Lys 510 505 500 Glu Glu Arg Arg Leu Thr Leu Ser Gly Lys Leu Ser Pro Glu Val Lys 520 · 525 515

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Glu C	gt cc ys Pr	a ccg o Pro	tgg Trp	gtt Val	gat Asp 135	agt Ser	atg Met	cgg Arg	agg Arg	agc Ser 140	tac Tyr	gtc Val	gga Gly	gat Asp	43	2

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Val Val His Gly Gly Glu Lys Ser Met Glu Glu Leu Asn Phe Ser 50 55

Asp Ser Asp Lys Glu Ser Thr Gly Cys Gln Ser Leu Pro Ala Thr Pro 65 70 75 80

Pro Arg Arg Arg Arg Gly Gly Gly Gly Gly Tyr Leu Ala 85

Val Ser Ser Pro Val Ser Gly Asp Lys Ala Tyr Ala Ser Glu Asn Glu 105 110 100

Val Gln Lys Thr Asn Asn Asn Gln Arg Arg Arg Arg Leu Lys Pro 115 Glu Cys Pro Pro Trp Val Asp Ser Met Arg Arg Ser Tyr Val Gly Asp 135 130 Glu Gln Ser Ser His Gly Gly Tyr Gly Gly Gly Val Val Val Thr 155 145 150 Arg Pro Ile Gly Gly Gly Arg Pro Leu Cys Met Asp Leu Glu Glu Val 170 165 Lys Ala Cys Lys Asp Leu Gly Phe Glu Leu Glu Pro Gly Arg Val Ser 185 180 Tyr Ser Gly Ser Thr Val Asp Thr Ser Ser Gly Gly Asn Ser Pro Ile 200 195 Ser Ser Asn His Arg Ile Ser Ser Pro Gly 215 210 <210> 95 <211> 2220 <212> DNA <213> Arabidopsis thaliana <220> <221> CDS <222> (1)..(2220) <223> ET3546 <400> 95 atg gaa gct atg ctt gtg gac tgt gta aac aac agt ctt cgt cat ttt 48 Met Glu Ala Met Leu Val Asp Cys Val Asn Asn Ser Leu Arg His Phe 10 5 gtc tac aaa aat gct att ttc atg tgc gag cgt ctc tgc gct gag ttt 96 Val Tyr Lys Asn Ala Ile Phe Met Cys Glu Arg Leu Cys Ala Glu Phe 30 20 cct tct gag gtt aat ttg cag cta tta gcc acc agc tac ctg cag aat 144

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Ile Pro Asn Gly Ala Ala Gly His Tyr Leu Leu Gly Leu Ile Tyr Lys
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Tyr Met Gln Gln Leu Ser Thr Ser Leu Gly Leu Asn Thr Tyr Asn Glu 165 170 175

Glu Arg Asn Ser Thr Ser Thr Lys Asn Thr Ser Ser Glu Asp Tyr Ser 180 185 190

Pro Arg Gln Ser Lys His Thr Gln Ser His Gly Leu Lys Asp Ile Ser Gly Asn Phe His Ser His Gly Val Asn Gly Gly Val Ser Asn Met Ser Phe Tyr Asn Thr Pro Ser Pro Val Ala Ala Gln Leu Ser Gly Ile Ala Pro Pro Pro Leu Phe Arg Asn Phe Gln Pro Ala Val Ala Asn Pro Asn Ser Leu Ile Thr Asp Ser Ser Pro Lys Ser Thr Val Asn Ser Thr Leu Gln Ala Pro Arg Arg Lys Phe Val Asp Glu Gly Lys Leu Arg Lys Ile Ser Gly Arg Leu Phe Ser Asp Ser Gly Pro Arg Arg Ser Ser Arg Leu Ser Ala Asp Ser Gly Ala Asn Ile Asn Ser Ser Val Ala Thr Val Ser Gly Asn Val Asn Asn Ala Ser Lys Tyr Leu Gly Gly Ser Lys Leu Ser Ser Leu Ala Leu Arg Ser Val Thr Leu Arg Lys Gly His Ser Trp Ala Asn Glu Asn Met Asp Glu Gly Val Arg Gly Glu Pro Phe Asp Asp Ser Arg Pro Asn Thr Ala Ser Thr Thr Gly Ser Met Ala Ser Asn Asp Gln Glu Asp Glu Thr Met Ser Ile Gly Gly Ile Ala Met Ser Ser Gln Thr Ile Thr Ile Gly Val Ser Glu Ile Leu Asn Leu Leu Arg Thr Leu Gly Glu Gly Cys Arg Leu Ser Tyr Met Tyr Arg Cys Gln Glu Ala Leu Asp

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Lys Leu Ser Tyr Leu Ala Gln Glu Leu Ile Ser Thr Asp Arg Leu Ala 500 505 510

Pro Gln Ser Trp Cys Ala Met Gly Asn Cys Tyr Ser Leu Gln Lys Asp 515 520 525

His Glu Thr Ala Leu Lys Asn Phe Leu Arg Ala Val Gln Leu Asn Pro 530 540

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Leu His Ala Leu Lys Arg Ser Glu Glu Ala Leu Glu Ile Met Glu Gln 625 630 635 640

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